



INTERNATIONAL APPLICATION PUBLISHED UNDER THE PATENT COOPERATION TREATY (PCT)

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(21) International Application Number: PCT/EP99/10209 (22) International Filing Date: 16 December 1999 (16.12.1999) (30) Priority Data: 98204291.3 16 December 1998 (16.12.1998) EP (60) Parent Application or Grant UNIVERSITY OF LIEGE [/]; (). MELICA HB [/]; (). SEGHERSGENTEC N.V. [/]; (). ANDERSSON, Leif [/]; (). GEORGES, Michel [/]; (). SPINCEMAILLE, Geert [/]; (). NEZER, Carine, Danielle, Andrée [/]; (). ANDERSSON, Leif [/]; (). GEORGES, Michel [/]; (). SPINCEMAILLE, Geert [/]; (). NEZER, Carine, Danielle, Andrée [/]; (). OTTEVANGERS, S., U.; ().		Published
(54) Title: SELECTING ANIMALS FOR PARENTALLY IMPRINTED TRAITS (54) Titre: SELECTION D'ANIMAUX EN FONCTION DE TRAITS COMMUNIQUEES PAR LEURS PARENTS (57) Abstract The invention relates to methods to select breeding animals or animals destined for slaughter for having desired genotypic or potential phenotypic properties, in particular related to muscle mass and/or fat deposition. The invention provides a method for selecting a pig for having desired genotypic or potential phenotypic properties comprising testing a sample from said pig for the presence of a quantitative trait locus (QTL) located at a Sus scrofa chromosome 2 mapping at position 2p1.7. (57) Abrégé L'invention concerne des procédés de sélection d'animaux reproducteurs ou destinés à l'abattoir sur la base des propriétés génotypiques désirées ou des propriétés phénotypiques potentielles qui sont notamment liées à la masse musculaire et/ou aux dépôts de lard. L'invention se rapporte à un procédé pour sélectionner un porc possédant des propriétés génotypiques désirées ou des propriétés phénotypiques potentielles, ledit procédé consistant à tester un échantillon provenant dudit porc pour vérifier la présence d'un locus quantitatif (QTL) présent dans la cartographie de chromosome 2 de Sus scrofa en position 2p1.7.		

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(54) Title: SELECTING ANIMALS FOR PARENTALLY IMPRINTED TRAITS			
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INTERNATIONAL SEARCH REPORT

International Application No.
PCT/EP 99/10209

A. CLASSIFICATION OF SUBJECT MATTER IPC 7 C12Q1/68 C07K14/65 A01K67/02		
According to International Patent Classification (IPC) or to both national classification and IPC		
B. FIELDS SEARCHED Minimum documentation searched (classification system followed by classification symbols) IPC 7 C12Q		
Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched		
Electronic data base consulted during the international search (name of data base and, where practical, search terms used) EPO-Internal, WPI Data, PAJ, MEDLINE, CHEM ABS Data, EMBASE, BIOSIS		
C. DOCUMENTS CONSIDERED TO BE RELEVANT		
Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	ANDERSSON-EKLUND ET AL.: "MAPPING QUANTITATIVE LOCI FOR CARCASS AND MEAT QUALITY TRAITS IN A WILD BOAR x LARGE WHITE INTERCROSS" J. ANIM. SCI., vol. 76, 1998, pages 694-700, XP002104406 cited in the application	1-3, 10-12
Y	See page 696, "Carcass Composition" and page 698, Fig. 1b. the whole document --- -/--	4-9, 13-27
<input checked="" type="checkbox"/> Further documents are listed in the continuation of box C. <input checked="" type="checkbox"/> Patent family members are listed in annex.		
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Date of the actual completion of the international search 1 August 2000		Date of mailing of the international search report 08/08/2000
Name and mailing address of the ISA European Patent Office, P.B. 5816 Paternlaan 2 NL - 2280 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl Fax: (+31-70) 340-3016		Authorized officer: Hagenmaier, S

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Int. Patent Application No.
PCT/EP 99/10209

C.(Continuation) DOCUMENTS CONSIDERED TO BE RELEVANT

Category	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	KOVACS AND KLÖTING: "MAPPING OF QUANTITATIVE TRAIT LOCI FOR BODY WEIGHT ON CHROMOSOMS 1 AND 4 IN THE RAT" BIOCHEMISTRY AND MOLECULAR BIOLOGY INTERNATIONAL, vol. 44, no. 2, February 1998 (1998-02), pages 399-405, XP002104407	1,2,10, 11
Y	the whole document	4-9, 13-27
Y	JOHANSSON ET AL.: "COMPARATIVE MAPPING REVEALS EXTENSIVE LINKAGE CONSERVATION-BUT WITH GENE ORDER REARRANGEMENTS-BETWEEN THE PIG AND THE HUMAN GENOMES" GENOMICS, vol. 25, 1995, pages 682-690, XP000610181 See Fig.1, pig chromosome 2 the whole document	4-9, 13-27
Y	REIK W ET AL: "IMPRINTING IN CLUSTERS: LESSONS FROM BECKWITH-WIEDEMANN SYNDROME" TRENDS IN GENETICS, vol. 13, no. 8, 1 August 1997 (1997-08-01), page 330-334 XP004084608 Igf2 the whole document	4-9, 13-27
Y	CATCHPOLE AND ENGSTRÖM: "NUCLEOTIDE SEQUENCE OF A PORCINE INSULINE-LIKE GROWTH FACTOR II cDNA" NUCLEIC ACIDS RESEARCH, vol. 18, no. 21, 1990, page 6430 XP002104409 cited in the application the whole document	15
A	ANDERSSON L ET AL: "GENETIC MAPPING OF QUANTITATIVE TRAIT LOCI FOR GROWTH AND FATNESS IN PIGS" SCIENCE, vol. 263, 25 March 1994 (1994-03-25), pages 1771-1774, XP002018359 cited in the application the whole document	
A	KNOTT ET AL.: "MULTIPLE MARKER MAPPING OF QUANTITATIVE TRAIT LOCI IN A CROSS BETWEEN OUTBRED WILD BOAR AND LARGE WHITE PIGS" GENETICS, vol. 149, June 1998 (1998-06), pages 1069-1080, XP002104410 cited in the application the whole document	
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INTERNATIONAL SEARCH REPORT

Int. Patent Application No.
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C. (Continuation) DOCUMENTS CONSIDERED TO BE RELEVANT

Category	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
A	WO 98 03682 A (UNIV IOWA RES FOUND) 29 January 1998 (1998-01-29) the whole document ----	
P, X	JEON ET AL.: "A PATERNALLY EXPRESSED QTL AFFECTING SKELETAL AND CARDIAC MUSCLE MASS IN PIGS MAPS TO THE IGF2 LOCUS" NAT.GENET., vol. 21, February 1999 (1999-02), pages 157-158, XP002104411 the whole document ----	1-27
P, X	NEZER ET AL.: "AN IMPRINTED QTL WITH MAJOR EFFECT ON MUSCLE MASS AND FAT DEPOSITION MAPS TO THE IGF2 LOCUS IN PIGS" NAT.GENET., vol. 21, February 1999 (1999-02), pages 155-156, XP002104412 the whole document -----	1-27

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INTERNATIONAL SEARCH REPORT

Information on patent family members

International Application No

PCT/EP 99/10209

Patent document cited in search report	Publication date	Patent family member(s)	Publication date
WO 9803682 A	29-01-1998	US 5935784 A	10-08-1999
		AU 3513297 A	10-02-1998
		BR 9710875 A	11-01-2000
		CN 1230227 A	29-09-1999
		CZ 9900161 A	16-06-1999
		EP 0958376 A	24-11-1999
		PL 331353 A	05-07-1999
		US 5939264 A	17-08-1999



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Description

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Title: Selecting animals for parentally imprinted traits.

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The invention relates to methods to select breeding animals or animals destined for slaughter for having desired genotypic or potential phenotypic properties, in particular related to muscle mass and/or fat deposition. Breeding schemes for domestic animals have so far focused on farm performance traits and carcass quality. This has resulted in substantial improvements in traits like reproductive success, milk production, lean/fat ratio, prolificacy, growth rate and feed efficiency. Relatively simple performance test data have been the basis for these improvements, and selected traits were assumed to be influenced by a large number of genes, each of small effect (the infinitesimal gene model). There are now some important changes occurring in this area. First, the breeding goal of some breeding organisations has begun to include meat quality attributes in addition to the "traditional" production traits. Secondly, evidence is accumulating that current and new breeding goal traits may involve relatively large effects (known as major genes), as opposed to the infinitesimal model that has been relied on so far.

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Modern DNA-technologies provide the opportunity to exploit these major genes, and this approach is a very promising route for the improvement of meat quality, especially since direct meat quality assessment is not viable for potential breeding animals. Also for other traits such as lean/fat ratio, growth rate and feed efficiency, modern DNA technology can be very effective. Also these traits are not always easy to measure in the living animal.

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The evidence for several of the major genes originally obtained using segregation analysis, i.e. without any DNA marker information. Afterwards molecular studies were performed to detect the location of these

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genes on the genetic map. In practice, and except for alleles of very large effect, DNA studies are required to dissect the genetic nature of most traits of economic importance. DNA markers can be used to localise genes or alleles responsible for qualitative traits like coat colour, and they can also be used to detect genes or alleles with substantial effects on quantitative traits like growth rate, IMF etc. In this case the approach is referred to as QTL (quantitative trait locus) mapping, wherein a QTL comprises at least a part of the nucleic acid genome of an animal where genetic information capable of influencing said quantitative trait (in said animal or in its offspring) is located. Information at DNA level can not only help to fix a specific major gene in a population, but also assist in the selection of a quantitative trait which is already selected for. Molecular information in addition to phenotypic data can increase the accuracy of selection and therefore the selection response.

Improving meat quality or carcass quality is not just about changing levels of traits like tenderness or marbling, but it is also about increasing uniformity. The existence of major genes provides excellent opportunities for improving meat quality because it allows large steps to be made in the desired direction. Secondly, it will help to reduce variation, since we can fix relevant genes in our products. Another aspect is that selecting for major genes allows differentiation for specific markets. Studies are underway in several species, particularly, pigs, sheep, deer and beef cattle.

In particular, intense selection for meat production has resulted in animals with extreme muscularity and leanness in several livestock species. In recent years it has become feasible to map and clone several of the genes causing these phenotypes, paving the way towards more efficient marker assisted selection, targeted drug development (performance enhancing products) and transgenesis. Mutations in the ryanodine receptor (Fuji

5 et al, 1991; MacLennan and Phillips, 1993) and myostatin
(Grobet et al, 1997; Kambadur et al, 1997; McPherron and
10 Lee, 1997) have been shown to cause muscular
hypertrophies in pigs and cattle respectively, while
5 genes with major effects on muscularity and/or fat
deposition have for instance been mapped to pig
chromosome 4 (Andersson et al, 1994) and sheep chromosome
18 (Cockett et al, 1996).

15 However, although there have been successes in
10 identifying QTLs, the information is currently of limited
use within commercial breeding programmes. Many workers
in this field conclude that it is necessary to identify
20 the particular genes underlying the QTL. This is a
substantial task, as the QTL region is usually relatively
15 large and may contain many genes. Identification of the
relevant genes from the many that may be involved thus
25 remains a significant hurdle in farm animals.

The invention provides a method for selecting a
20 domestic animal for having desired genotypic or potential
phenotypic properties comprising testing said animal for
30 the presence of a parentally imprinted qualitative or
quantitative trait locus (QTL). Herein, a domestic animal
is defined as an animal being selected or having been
35 derived from an animal having been selected for having
25 desired genotypic or potential phenotypic properties.

Domestic animals provide a rich resource of genetic
and phenotypic variation, traditionally domestication
40 involves selecting an animal or its offspring for having
30 desired genotypic or potential phenotypic properties.
This selection process has in the past century been
45 facilitated by growing understanding and utilisation of
the laws of Mendelian inheritance. One of the major
problems in breeding programs of domestic animals is the
35 negative genetic correlation between reproductive
capacity and production traits. This is for example the
50 case in cattle (a high milk production generally results

5 in slim cows and bulls) poultry, broiler lines have a low
level of egg production and layers have generally very
low muscle growth), pigs (very prolific sows are in
10 general fat and have comparatively less meat) or sheep
5 (high prolific breeds have low carcass quality and vice
versa). The invention now provides that knowledge of the
parental imprinting character of various traits allows to
select for example sire lines homozygous for a paternally
15 imprinted QTL for example linked with muscle production
or growth; the selection for such traits can thus be less
stringent in dam lines in favour of the reproductive
quality. The phenomenon of genetic or parental imprinting
20 has never been utilised in selecting domestic animals, it
was never considered feasible to employ this elusive
genetic characteristic in practical breeding programmes.
The invention provides a breeding programme, wherein
25 knowledge of the parental imprinting character of a
desired trait, as demonstrated herein, results in a
breeding programme, for example in a BLUP programme, with
a modified animal model. This increases the accuracy of
30 the breeding value estimation and speeds up selection
compared to conventional breeding programmes. Until now,
the effect of a parentally imprinted trait in the
estimation of a conventional BLUP programme was
35 neglected; using and understanding the parental character
of the desired trait, as provided by the invention,
allows selecting on parental imprinting, even without DNA
testing. For example, selecting genes characterised by
40 paternal imprinting is provided to help increase
uniformity; a (terminal) parent homozygous for the "good
or wanted" alleles will pass them to all offspring,
45 regardless of the other parent's alleles, and the
offspring will all express the desired parent's alleles.
This results in more uniform offspring. Alleles that are
35 interesting or favourable from the maternal side or often
the ones that have opposite effects to alleles from the
paternal side. For example, in meat animals such as pigs
50 alleles linked with meat quality traits such as intra-

muscular fat or muscle mass could be fixed in the dam lines while alleles linked with reduced back fat could be fixed in the sire lines. Other desirable combinations are for example fertility and/or milk yield in the female line with growth rates and/or muscle mass in the male lines.

In a preferred embodiment, the invention provides a method for selecting a domestic animal for having desired genotypic or potential phenotypic properties comprising testing a nucleic acid sample from said animal for the presence of a parentally imprinted quantitative trait locus (QTL). A nucleic acid sample can in general be obtained from various parts of the animal's body by methods known in the art. Traditional samples for the purpose of nucleic acid testing are blood samples or skin or mucosal surface samples, but samples from other tissues can be used as well, in particular sperm samples, oocyte or embryo samples can be used. In such a sample, the presence and/or sequence of a specific nucleic acid, be it DNA or RNA, can be determined with methods known in the art, such as hybridisation or nucleic acid amplification or sequencing techniques known in the art. The invention provides testing such a sample for the presence of nucleic acid wherein a QTL or allele associated therewith is associated with the phenomenon of parental imprinting, for example where it is determined whether a paternal or maternal allele of said QTL is capable of being predominantly expressed in said animal.

The purpose of breeding programs in livestock is to enhance the performances of animals by improving their genetic composition. In essence this improvement accrues by increasing the frequency of the most favourable alleles for the genes influencing the performance characteristics of interest. These genes are referred to as QTL. Until the beginning of the nineties, genetic improvement was achieved via the use of biometrical methods, but without molecular knowledge of the underlying QTL.

5 Since the beginning of the nineties and due to
recent developments in genomics, it is conceivable to
identify the QTL underlying a trait of interest. The
invention now provides identifying and using parentally
10 5 imprinted QTLs which are useful for selecting animals by
mapping quantitative trait loci. Again, the phenomenon of
genetic or paternal imprinting has never been utilised in
selecting domestic animals, it was never considered
15 feasible to employ this elusive genetic characteristic in
practical breeding programmes. For example Kovacs and
10 Kloting (Biochem. Mol. Biol. Int. 44:399-405, 1998),
where parental imprinting is not mentioned, and not
20 suggested, found linkage of a trait in female rats, but
not in males, suggesting a possible sex specificity
15 associated with a chromosomal region, which of course
excludes parental imprinting, a phenomenon wherein the
imprinted trait of one parent is preferably but gender-
25 aspecifically expressed in his or her offspring.

The invention provides the initial localisation of a
20 parentally imprinted QTL on the genome by linkage
analysis with genetic markers, and the actual
30 identification of the parentally imprinted gene(s) and
causal mutations therein. Molecular knowledge of such a
parentally imprinted QTL allows for more efficient
25 breeding designs herewith provided. Applications of
molecular knowledge of parentally imprinted QTLs in
breeding programs include: marker assisted segregation
analysis to identify the segregation of functionally
distinct parentally imprinted QTL alleles in the
40 30 populations of interest, marker assisted selection (MAS)
performed within lines to enhance genetic response by
increasing selection accuracy, selection intensity or by
reducing the generation interval using the understanding
45 of the phenomenon of parental imprinting, marker assisted
35 introgression (MAI) to efficiently transfer favourable
parentally imprinted QTL alleles from a donor to a
recipient population, genetic engineering of the
50 identified parentally QTL and genetic modification of the
breeding stock using transgenic technology, development

5 of performance enhancing products using targeted drug development exploiting molecular knowledge of said QTL.

The inventors undertook two independent experiments to determine the practical use of parental imprinting of
10 5 a QTL.

In a first experiment, performed in a previously described Piétrain x Large White intercross, the likelihood of the data were computed under a model of paternal (paternal allele only expressed) and maternal
15 10 imprinting (maternal allele only expressed) and compared with the likelihood of the data under a model of a conventional "Mendelian" QTL. The results strikingly demonstrated that the QTL was indeed paternally
20 expressed, the QTL allele (Piétrain or Large White) inherited from the F₁ sow having no effect whatsoever on the carcass quality and quantity of the F₂ offspring. It was seen that very significant lodscores were obtained
25 when testing for the presence of a paternally expressed QTL, while there was no evidence at all for the segregation of a QTL when studying the chromosomes
30 transmitted by the sows. The same tendency was observed for all traits showing that the same imprinted gene is responsible for the effects observed on the different traits. Table 1 reports the maximum likelihood (ML) phenotypic means for the F₂ offspring sorted by inherited
35 paternal QTL allele.

In a second experiment performed in the Wild Boar X Large White intercross, QTL analyses of body composition, fatness, meat quality, and growth traits was carried out
40 30 with the chromosome 2 map using a statistical model testing for the presence of an imprinting effect. Clear evidence for a paternally expressed QTL located at the very distal tip of 2p was obtained (Fig. 2; Table1). The clear paternal expression of a QTL is illustrated by the
45 35 least squares means which fall into two classes following the population origin of the paternally inherited allele (Table 1). For a given paternally imprinted QTL,
50 implementation of marker assisted segregation analysis, selection (MAS) and introgression (MAI), can be performed

5 using genetic markers that are linked to the QTL, genetic markers that are in linkage disequilibrium with the QTL, or using the actual causal mutations within the QTL.

Understanding the parent-of-origin effect

10 5 characterising a QTL allows for its optimal use in breeding programs. Indeed, marker assisted segregation analysis under a model of parental imprinting will yield better estimates of QTL allele effects. Moreover it
15 10 allows for the application of specific breeding schemes to optimally exploit a QTL. In one embodiment of the invention, the most favourable QTL alleles would be fixed in breeding animal lines and for example used to generate commercial, crossbred males by marker assisted selection
20 (MAS, within lines) and marker assisted introgression (MAI, between lines). In another embodiment, the worst QTL alleles would be fixed in the animal lines used to generate commercial crossbred females by MAS (within
25 lines) and MAI (between lines).

In a preferred embodiment of the invention, said
20 animal is a pig. Note for example that the invention provides the insight that today half of the offspring from commercially popular Piétrain, Large White crossbred
30 boars inherit an unfavourable Large White muscle mass QTL as provided by the invention causing considerable loss, and the invention now for example provides the
35 possibility to select the better half of the population in that respect. However, it is also possible to select commercial sow lines enriched with the in the boars unfavourable alleles, allowing to equip the sows with
40 30 other alleles more desirable for for example reproductive purposes.

In a preferred embodiment of a method provided by the invention, said QTL is located at a position
45 corresponding to a QTL located at chromosome 2 in the pig. For example, it is known from comparative mapping data between pig and human, including bidirectional chromosome painting, that SSC2p is homologous to
50 35 HSA11pter-q13^{11,12}. HSA11pter-q13 is known to harbour a

5 cluster of imprinted genes: IGF2, INS2, H19, MAH2, P57^{KIP2},
K_vLQTL1, Tapal/CD81, Orctl2, Impt1 and Ipl. The cluster
of imprinted genes located in HSA11pter-q13 is
characterised by 8 maternally expressed genes H19, MASH2,
10 5 P57^{KIP2}, K_vLQTL1, TAPAL/CD81, ORCTL2, IMPT1 and IP1, and
two paternally expressed genes: IGF2 and INS. However,
Johanson et al (Genomics 25:682-690, 1995) and Reik et al
(Trends in Genetics, 13:330-334, 1997) show that the
15 whereabouts of these loci in various animals are not
20 clear. For example, the HSA11 and MMU7 loci do not
correspond among each other, the MMU7 and the SSC2 loci
do not correspond, whereas the HSA11 and SSC2 loci seem
to correspond, and no guidance is given where one or more
of for example the above identified parentally expressed
15 individual genes are localised on the three species'
chromosomes.

25 Other domestic animals, such as cattle, sheep,
poultry and fish, having similar regions in their genome
harbouring such a cluster of imprinted genes or QTLs, the
20 invention herewith provides use of these orthologous
regions of other domestic animals in applying the
phenomenon of parental imprinting in breeding programmes.
In pigs, said cluster is mapped at around position 2p1.7
of chromosome 2, however, a method as provided by the
35 invention employing (fragments of) said maternally or
paternally expressed orthologous or homologous genes or
QTLs are advantageously used in other animals as well for
breeding and selecting purposes. For example, a method is
provided wherein said QTL is related to the potential
40 30 muscle mass and/or fat deposition, preferably with
limited effects on other traits such as meat quality and
daily gain of said animal or wherein said QTL comprises
at least a part of an insulin-like growth factor-2 (IGF2)
45 allele. Reik et al (Trends in Genetics, 13:330-334, 1997)
35 explain that this gene in humans is related to Beckwith-
Wiedemann syndrome, an apparently parentally imprinted
disease syndrome most commonly seen with human foetuses,
50 where the gene has an important role in prenatal

5 development. No relationship is shown or suggested with postnatal development relating to muscle development or fatness in (domestic) animals.

10 In a preferred embodiment, the invention provides a method for selecting a pig for having desired genotypic or potential phenotypic properties comprising testing a sample from said pig for the presence of a quantitative trait locus (QTL) located at a *Sus scrofa* chromosome 2 mapping at position 2p1.7. In particular, the invention 15 relates to the use of genetic markers for the telomeric end of pig chromosome 2p in marker selection (MAS) of a parentally imprinted Quantitative Trait Locus (QTL) affecting carcass yield and quality in pigs. Furthermore, the invention relates to the use of genetic markers 20 associated with the IGF2 locus in MAS in pigs, such as polymorphisms and microsatellites and other characterising nucleic acid sequences shown herein, such as shown in figures 4 to 10. In a preferred embodiment, the invention provides a QTL located at the distal tip of *Sus scrofa* 25 chromosomes 2 with effects on varies measurements of carcass quality and quantity, particularly muscle mass and fat deposition.

30 In a first experiment, a QTL mapping analysis was performed in a Wild Boar X Large White intercross counting 200 F₂ individuals. The F₂ animals were 35 sacrificed at a live eight of at least 80 kg or at a maximum age of 190 days. Phenotypic data on birth weight, growth, fat deposition, body composition, weight of internal organs, and meat quality were collected; a 40 detailed description of the phenotypic traits are provided by Andersson et al¹ and Andersson-Eklund et al¹.

45 A QTL (without any significant effect on back-fat thickness) at an unspecified locus on the proximal end of chromosome 2 with moderate effect on muscle mass, and 35 located about 30cM away from the parentally imprinted QTL reported here, was previously reported by the inventors; whereas the QTL as now provided has a very large effect, 50 explaining at least 20-30% of variance, making the QTL of

the present invention commercially very attractive, which is even more so because the present QTL is parentally imprinted. The marker map of chromosome 2p was improved as part of this invention by adding microsatellite

markers in order to cover the entire chromosome arm. The following microsatellite markers were used: Swc9, Sw2443, Sw2623, and Swr2516, all from the distal end of 2p⁷. QTL analyses of body composition, fatness, meat quality, and growth traits were carried out with the new chromosome 2 map. Clear evidence for a QTL located at the very distal tip of 2p was obtained (Fig. 1; Table 1). The QTL had very large effects on lean meat content in ham and explained an astonishing 30% of the residual phenotypic variance in the F₂ population. Large effects on the area of the longissimus dorsi muscle, on the weight of the heart, and on back-fat thickness (subcutaneous fat) were also noted. A moderate effect on one meat quality trait, reflectance value, was indicated. The QTL had no significant effect on abdominal fat, birth weight, growth, weight of liver, kidney, or spleen (data not shown). The Large White allele at this QTL was associated with larger muscle mass and reduced back-fat thickness consistent with the difference between this breed and the Wild Boar population.

In a second experiment, QTL mapping was performed in a Piétrain X Large White intercross comprising 1125 F₂ offspring. The Large White and Piétrain parental breeds differ for a number of economically important phenotypes. Piétrains are famous for their exceptional muscularity and leanness¹⁰ (Figure 2, while Large Whites show superior growth performance. Twenty-one distinct phenotypes measuring growth performance (5), muscularity (6), fat deposition (6), and meat quality (4), were recorded on all F₂ offspring. In order to map QTL underlying the genetic differences between these breeds, the inventors undertook a whole genome scan using microsatellite markers on an initial sample of 677 F₂ individuals. The following microsatellite marker map was used to analyse

5 chromosome 2::SW2443, SWC9 and SW2623, SWR2516-(0,20)-
SWR783-(0,29)-SW240-(0,20)-SW776-(0,08)-S0010-(0,04)-
SW1695-(0,36)-SWR308. Analysis of pig chromosome 2 using
10 a Maximum Likelihood multipoint algorithm, revealed
5 highly significant lodscores (up to 20) for three of the
six phenotypes measuring muscularity (% lean cuts, % ham,
% loin) and three of the six phenotypes measuring fat
deposition (back-fat thickness (BFT), % backfat, % fat
15 cuts) at the distal end of the short arm of chromosome 2
10 (Figure 1). Positive lodscores were obtained in the
corresponding chromosome region for the remaining six
muscularity and fatness phenotypes, however, not reaching
the experiment-wise significance threshold ($\alpha=5\%$). There
20 was no evidence for an effect of the corresponding QTL on
15 growth performance (including birth weight) or recorded
meat quality measurements (data not shown). To confirm
this finding, the remaining sample of 355 F₂ offspring was
25 genotyped for the four most distal 2p markers and QTL
analysis performed for the traits yielding the highest
20 lodscores in the first analysis. Lodscores ranged from
2.1 to 7.7, clearly confirming the presence of a major
30 QTL in this region. Table 2 reports the corresponding ML
estimates for the three genotypic means as well as the
residual variance. Evidence based on marker assisted
35 segregation analysis points towards residual segregation
at this locus within the Piétrain population.

These experiments therefore clearly indicated
the existence of a QTL with major effect on carcass
quality and quantity on the telomeric end of pig
40 chromosome arm 2p; the likely existence of an allelic
30 series at this QTL with at least three alleles: Wild-Boar
< Large White < Piétrain, and possibly more given the
observed segregation within the Piétrain breed.

45 The effects of the identified QTL on muscle mass and
35 fat deposition are truly major, being of the same
magnitude of those reported for the CRC locus though
apparently without the associated deleterious effects on
50 meat quality. We estimate that both loci jointly explain

close to 50% of the Piétrain versus Large White breed difference for muscularity and leanness. The QTL had very large effects on lean meat content in ham and explained an astonishing 30% of the residual phenotypic variance in the F₂ population. Large effects on the area of the longissimus dorsi muscle, on the weight of the heart, and on back-fat thickness (subcutaneous fat) were also noted. A moderate effect on one meat quality trait, reflectance value, was indicated. The QTL had no significant effect on abdominal fat, birth weight, growth, weight of liver, kidney, or spleen (data not shown). The Large White allele at this QTL, when compared to the Wild Boar allele, was associated with larger muscle mass and reduced back-fat thickness consistent with the difference between this breed and the Wild Boar population. The strong imprinting effect observed for all affected traits shows that a single causative locus is involved. The pleiotropic effects on skeletal muscle mass and the size of the heart appear adaptive from a physiological point of view as a larger muscle mass requires a larger cardiac output.

In a further embodiment, the invention provides a method for selecting a pig for having desired genotypic or potential phenotypic properties comprising testing a sample from said pig for the presence of a quantitative trait locus (QTL) located at a *Sus scrofa* chromosome 2 mapping at position 2p1.7., wherein said QTL comprises at least a part of a *Sus scrofa* insulin-like growth factor-2 (IGF2) allele or a genomic area closely related thereto, such as polymorphisms and microsatellites and other characterising nucleic acid sequences shown herein, such as shown in figures 4 to 10. The important role of IGF2 for prenatal development is well-documented from knock-out mice as well as from its causative role in the human Beckwith-Wiedemann syndrome. This invention demonstrates an important role for the IGF2-region also for postnatal development.

5 To show the role of Igf2 the inventors performed the following three experiments:

10 A genomic IGF2 clone was isolated by screening a porcine BAC library. FISH analysis with this BAC clone
5 gave a strong consistent signal on the terminal part of chromosome 2p.

15 A polymorphic microsatellite is located in the 3'UTR of IGF2 in mice (GenBank U71085), humans (GenBank S62623), and horse (GenBank AF020598). The possible
10 presence of a corresponding porcine microsatellite was investigated by direct sequencing of the IGF2 3'UTR using the BAC clone. A complex microsatellite was identified
20 about 800bp downstream of the stop codon; a sequence comparison revealed that this microsatellite was
15 identical to a previously described anonymous microsatellite, Swc9⁶. This marker was used in the initial
25 QTL mapping experiments and its location on the genetic map correspond with the most likely position of the QTL both in the Piétrain X Large White and in the Large White
20 x Wild Boar pedigree.

30 Analysis of skeletal muscle and liver cDNA from 10-week old fetuses heterozygous for a nt241 (G-A) transversion in the second exon of the porcine IGFII gene and SWC9, shows that the IGFII gene is imprinted in these
25 tissues in the pig as well and only expressed from the paternal allele.

35 Based on a published porcine adult liver cDNA sequence¹⁶, the inventors designed primer pairs allowing to amplify the entire IgfII coding sequence with 222 bp
40 of leader and 280 bp of trailer sequence from adult skeletal muscle cDNA. Piétrain and Large White RT-PCR products were sequenced indication that the coding
45 sequences are identical in both breeds and with the published sequence. However, a G→A transition was found
35 in the leader sequence corresponding to exon 2 in man. Following conventional nomenclature, this polymorphism will be referred to as nt241(G-A). We developed a
50 screening test for this single nucleotide polymorphism

9 (SNP) based on the ligation amplification reaction (LAR), allowing us to genotype our pedigree material. Based on these data, *IgfII* was shown to colocalize with the SWC9 microsatellite marker ($\theta=0\%$), therefore

virtually coinciding with the most likely position of the QTL, and well within the 95% support interval for the QTL. Subsequent sequence analysis demonstrated that the microsatellite marker SWC9 is actually located within the 3'UTR of the *IgfII* gene.

As previously mentioned, the knowledge of this QTL provides a method for the selection of animals such as pigs with improved carcass merit. Different embodiments of the invention are envisaged, including: marker assisted segregation analysis to identify the segregation of functionally distinct QTL alleles in the populations of interest; marker assisted selection (MAS) performed within lines to enhance genetic response by increasing selection accuracy, selection intensity or by reducing the generation interval; marker assisted introgression (MAI) to efficiently transfer favourable QTL alleles from a donor to a recipient population; thereby enhancing genetic response in the recipient population. Implementation of embodiments marker assisted segregation analysis, selection (MAS) and introgression (MAI), can be performed using genetic markers that are linked to the QTL; genetic markers that are in linkage disequilibrium with the QTL, the actual causal mutations within the QTL.

In a further embodiment, the invention provides a method for selecting a pig for having desired genotypic or potential phenotypic properties comprising testing a sample from said pig for the presence of a quantitative trait locus (QTL) located at a *Sus scrofa* chromosome 2 mapping at position 2p1.7., wherein said QTL is paternally expressed, i.e. is expressed from the paternal allele. In man and mouse, *Igf2* is known to be imprinted and to be expressed exclusively from the paternal allele in several tissues. Analysis of skeletal muscle cDNA from

5 pigs heterozygous for the SNP and/or SWC9, shows that the same imprinting holds in the pig as well. Understanding the parent-of-origin effect characterising the QTL as provided by the invention now allows for its optimal use
10 5 in breeding programs. Indeed, today half of the offspring from commercially popular Piétrain x Large White crossbred boars inherit the unfavourable Large White allele causing considerable loss. Using a method as
15 provide by the invention avoids this problem.

10 The invention furthermore provides an isolated and/or recombinant nucleic acid or functional fragment derived thereof comprising a parentally imprinted
20 quantitative trait locus (QTL) or fragment thereof capable of being predominantly expressed by one parental
15 allele. Having such a nucleic acid as provided by the invention available allows constructing transgenic
25 animals wherein favourable genes are capable of being exclusively or predominantly expressed by one parental allele, thereby equipping the offspring of said animal
20 homozygous for a desired trait with desired properties related to that parental allele that is expressed.

In a preferred embodiment, the invention provides an isolated and/or recombinant nucleic acid or fragment derived thereof comprising a synthetic parentally
35 25 imprinted quantitative trait locus (QTL) or functional fragment thereof derived from at least one chromosome. Synthetic herein describes a parentally expressed QTL wherein various elements are combined that originate from
40 distinct locations from the genome of one or more
30 animals. The invention provides recombinant nucleic acid wherein sequences related to parental imprinting of one QTL are combined with sequences relating to genes or
45 favourable alleles of a second QTL. Such a gene construct is favourably used to obtain transgenic animals wherein
35 the second QTL has been equipped with paternal imprinting, as opposed to the inheritance pattern in the native animal from which the second QTL is derived. Such
50 a second QTL can for example be derived from the same

5 chromosome where the parental imprinting region is
located, but can also be derived from a different
chromosome from the same or even a different species. In
the pig, such a second QTL can for example be related to
10 5 an oestrogen receptor (ESR)-gene (Rothschild et al, PNAS,
93, 201-201, 1996) or a FAT-QTL (Andersson, Science, 263,
1771-1774, 1994) for example derived from an other pig
chromosome, such as chromosome 4. A second or further QTL
15 can also be derived from another (domestic) animal or a
10 human.

The invention furthermore provides an isolated
and/or recombinant nucleic acid or functional fragment
20 derived thereof at least partly corresponding to a QTL of
a pig located at a *Sus scrofa* chromosome 2 mapping at
15 position 2p1.7 wherein said QTL is related to the
potential muscle mass and/or fat deposition of said pig
and/or wherein said QTL comprises at least a part of a
25 *Sus scrofa* insulin-like growth factor-2 (IGF2) allele,
preferably at least spanning a region between INS and
20 H19, or preferably derived from a domestic pig, such as a
Pietrain, Meishan, Duroc, Landrace or Large White, or
30 from a Wild Boar. For example, a genomic IGF2 clone was
isolated by screening a porcine BAC library. FISH
analysis with this BAC clone gave a strong consistent
35 25 signal on the terminal part of chromosome 2p. A
polymorphic microsatellite is located in the 3'UTR of
IGF2 in mice (GenBank U71085), humans (GenBank S62623),
and horse (GenBank AF020598). The possible presence of a
40 corresponding porcine microsatellite was investigated by
30 direct sequencing of the IGF2 3'UTR using the BAC clone.
A complex microsatellite was identified about 800 bp
downstream of the stop codon; a sequence comparison
45 revealed that this microsatellite is identical to a
previously described anonymous microsatellite, Swc9. PCR
35 25 primers were designed and the microsatellite (IGF2_{ms}) was
found to be highly polymorphic with three different
50 alleles among the two Wild Boar founders and another two

among the eight Large White founders. *IGF2ms* was fully informative in the intercross as the breed of origin as well as the parent of origin could be determined with confidence for each allele in each F_2 animal.

A linkage analysis using the intercross pedigree was carried out with *IGF2ms* and the microsatellites *Sw2443*, *Sw2623*, and *Swr2516*, all from the distal end of 2p⁷. *IGF2* was firmly assigned to 2p by highly significant lod scores (e.g. $Z=89.0$, $\theta=0.003$ against *Swr2516*). Multipoint analyses, including previously typed chromosome 2 markers, revealed the following order of loci (sex-average map distances in Kosambi cM): *Sw2443/Swr2516*-0.3-*IGF2*-14.9-*Sw2623*-10.3-*Sw256*. No recombinant was observed between *Sw2443* and *Swr2516*, and the suggested proximal location of *IGF2* in relation to these loci is based on a single recombinant giving a lod score support of 0.8 for the reported order. The most distal marker in our previous QTL study, *Sw256*, is located about 25 cM from the distal end of the linkage group.

The invention furthermore provides use of a nucleic acid or functional fragment derived thereof according to the invention in a method according to the invention. In a preferred embodiment, use of a method according to invention is provided to select a breeding animal or animal destined for slaughter, or embryos or semen derived from these animals for having desired genotypic or potential phenotypic properties. In particular, the invention provides such use wherein said properties are related to muscle mass and/or fat deposition. The QTL as provided by the invention may be exploited or used to improve for example lean meat content or back-fat thickness by marker assisted selection within populations or by marker assisted introgression of favorable alleles from one population to another. Examples of marker assisted selection using the QTL as provided by the invention are use of marker assisted segregation analysis

5 with linked markers or with markers in disequilibrium to
identify functionally distinct QTL alleles. Furthermore,
identification of a causative mutation in the QTL is now
possible, again leading to identify functionally distinct
10 5 QTL alleles. Such functionally distinct QTL alleles
located at the distal tip of chromosome 2p with large
effects on skeletal muscle mass, the size of the heart,
and on back-fat thickness are also provided by the
15 invention. The observation of a similar QTL effect in a
10 Large White x Wild Boar as well as in a Piétrain x Large
White intercross provides proof of the existence of a
series of at least three distinct functional alleles.
20 Moreover, preliminary evidence based on marker assisted
segregation analysis points towards residual segregation
15 at this locus within the Piétrain population (data not
shown). The occurrence of an allelic series as provided
by the invention allows identifying causal polymorphisms
which - based on the quantitative nature of the observed
effect - are unlikely to be gross gene alterations but
20 rather subtle regulatory mutations. The effects on muscle
mass of the three alleles rank in the same order as the
breeds in which they are found i.e. Piétrain pigs are
more muscular than Large White pigs that in turn have
30 higher lean meat content than Wild Boars. The invention
25 furthermore provides use of the alleles as provided by
the invention for within line selection or for marker
assisted introgression using linked markers, markers in
disequilibrium or alleles comprising causative mutations.

30 The invention furthermore provides an animal
selected by using a method according to the invention.
For example, a pig characterised in being homozygous for
45 an allele in a QTL located at a Sus scrofa chromosome 2
mapping at position 2p1.7 can now be selected and is thus
provided by the invention. Since said QTL is related to
35 the potential muscle mass and/or fat deposition of said
pig and/or said QTL comprises at least a part of a Sus
50 scrofa insulin-like growth factor-2 (IGF2) allele, it is

5 possible to select promising pigs to be used for breeding
or to be slaughtered. In particular an animal according
to the invention which is a male is provided. Such a
male, or its sperm or an embryo derived thereof can
10 advantageously be used in breeding animals for creating
breeding lines or for finally breeding animals destined
for slaughter. In a preferred embodiment of such use as
provided by the invention, a male, or its sperm,
15 deliberately selected for being homozygous for an allele
causing the extreme muscular hypertrophy and leanness,
20 is used to produce offspring heterozygous for such an
allele. Due to said allele's paternal expression, said
offspring will also show the favourable traits for
example related to muscle mass, even if the parent female
15 has a different genetic background. Moreover, it is now
possible to positively select the female(s) for having
different traits, for example related to fertility,
25 without having a negative effect on the muscle mass trait
that is inherited from the allele from the selected male.
20 For example, earlier such males could occasionally be
seen with Pietrain pigs but genetically it was not
30 understood how to most profitably use these traits in
breeding programmes.

Furthermore, the invention provides a transgenic
35 animal, sperm and an embryo derived thereof, comprising a
synthetic parentally imprinted QTL or functional fragment
thereof as provided by the invention, i.e. it is provided
by the invention to introduce a favourable recombinant
40 allele; for example introduce the oestrogen receptor
30 locus related to increased litter size of an animal
homozygously in a parentally imprinted region of a
grandparent animal (for example the father of a hybrid
sow if the region was paternally imprinted and the
45 grandparent was a boar); to introduce a favourable fat-
35 related allele or muscle mass-related recombinant allele
in a paternally imprinted region, and so on. Recombinant
alleles that are interesting or favourable from the
50 maternal side or often the ones that have opposite
effects to alleles from the paternal side. For example,

5 in meat animals such as pigs recombinant alleles linked
with meat quality traits such as intra-muscular fat or
muscle mass could be fixed in the dam lines while
10 recombinant alleles linked with reduced back fat could be
5 fixed in the sire lines. Other desirable combinations are
for example fertility and/or milk yield in the female
line with growth rates and/or muscle mass in the male
lines.

15 The invention is further explained in the detailed
10 description without limiting the invention.

Detailed description.

20 Example 1: Wild Boar x Large White intercrosses

15 Methods

25 Isolation of an IGF2 BAC clone and fluorescent *in situ*
hybridization (FISH). IGF2 primers (F:5'-
20 GGCAAGTTCTTCCGCTAATGA-3' and R:5'-GCACCGCAGAATTACGACAA-
30 3') for PCR amplification of a part of the last exon and
3'UTR were designed on the basis of a porcine IGF2 cDNA
sequence (GenBank X56094). The primers were used to
screen a porcine BAC library and the clone 253G10 was
35 25 isolated. Crude BAC DNA was prepared as described²⁴. The
BAC DNA was linearized with EcoRV and purified with
QIAEXII (QIAGEN GmbH, Germany). The clone was labeled
40 with biotin-14-dATP using the GIBCO-BRL Bionick labeling
system (BRL18246-015). Porcine metaphase chromosomes were
30 obtained from pokeweed (Seromed) stimulated lymphocytes
using standard techniques. The slides were aged for two
45 days at room temperature and then kept at -20°C until
use. FISH analysis was carried out as previously
described²⁵. The final concentration of the probe in the
50 35 hybridization mix was 10 ng/μl. Repetitive sequences were
suppressed with standard concentrations of porcine

5 genomic DNA. After post-hybridization washing, the
biotinylated probe was detected with two layers of
avidin-FITC (Vector A-2011). The chromosomes were
counterstained with 0.3 mg/ml DAPI (4,6-Diamino-2-
10 phenylindole; Sigma D9542), which produced a G-banding
like pattern. No posthybridization banding was needed,
since chromosome 2 is easily recognized without banding.
A total of 20 metaphase spreads were examined under an
15 Olympus BX-60 fluorescence microscope connected to an
20 IMAC-CCD S30 video camera and equipped with an ISIS 1.65
(Metasystems) software.

20 Sequence, microsatellite, and linkage analysis.

15 About two µg of linearized and purified BAC DNA was used
25 for direct sequencing with 20 pmoles of primers and
BigDye Terminator chemistry (Perkin Elmer, USA). DNA
sequencing was done from the 3' end of the last exon
towards the 3' end of the UTR until a microsatellite was
30 detected. A primer set (F:5'-GTTTCTCTGTACCCACACGCATCCC-
3' and R:5'-Fluorescein-CTACAAGCTGGGCTCAGGG-3') was
designed for the amplification of the *IGF2* microsatellite
which is about 250 bp long and located approximately 800
35 bp downstream from the stop codon. The microsatellite was
25 PCR amplified using fluorescently labeled primers and the
genotyping was carried out using an ABI377 sequencer and
the GeneScan/Genotyper softwares (Perkin Elmer, USA).
40 Two-point and multipoint linkage analysis were done with
the Cri-Map software²⁶.

30 Animals and phenotypic data.

45 The intercross pedigree comprised two European Wild Boar
males and eight Large White females, 4 F₁ males and 22 F₁
50 females, and 200 F₂ progeny¹. The F₂ animals were
sacrificed at a live weight of at least 80 kg or at a

5 maximum age of 190 days. Phenotypic data on birth weight,
growth, fat deposition, body composition, weight of
internal organs, and meat quality were collected; a
detailed description of the phenotypic traits are
10 5 provided by Andersson *et al.*¹ and Andersson-Eklund *et al.*⁴

15 Statistical analysis.

10 Interval mapping for the presence of QTL were carried out
with a least squares method developed for the analysis of
crosses between outbred lines²⁷. The method is based on
the assumption that the two divergent lines are fixed for
alternative QTL alleles. There are four possible
15 genotypes in the F_2 generation as regards the
grandparental origin of the alleles at each locus. This
makes it possible to fit three effects: additive,
dominance, and imprinting². The latter is estimated as
the difference between the two types of heterozygotes,
30 20 the one receiving the Wild Boar allele through an F_1 sire
and the one receiving it from an F_1 dam. An F-ratio was
calculated using this model (with 3 d.f.) versus a
reduced model without a QTL effect for each cM of
35 chromosome 2. The most likely position of a QTL was
obtained as the location giving the highest F-ratio.
Genome-wise significance thresholds were obtained
40 empirically by a permutation test²⁸ as described². The
QTL model including an imprinting effect was compared
with a model without imprinting (with 1 d.f.) to test
30 whether the imprinting effect was significant.

45 The statistical models also included the fixed
effects and covariates that were relevant for the
respective traits; see Andersson-Eklund *et al.*⁴ for a
more detailed description of the statistical models used.
50 35 Family was included to account for background genetic

5 effects and maternal effects. Carcass weight was included
as a covariate to discern QTL effects on correlated
traits, which means that all results concerning body
composition were compared at equal weights. Least-squares
10 5 means for each genotype class at the *IGF2* locus were
estimated with a single point analysis using Procedure
GLM of SAS²⁹; the model included the same fixed effects
and covariates as used in the interval mapping analyses.
15 The QTL shows a clear parent of origin-specific
20 10 expression and the map position coincides with that of
the insulin-like growth factor II gene (*IGF2*), indicating
IGF2 as the causative gene. A highly significant
segregation distortion (excess of Wild Boar-derived
alleles) was also observed at this locus. The results
25 15 demonstrate an important effect of the *IGF2* region on
postnatal development and it is possible that the
presence of a paternally expressed *IGF2*-linked QTL in
humans and in rodent model organisms has so far been
overlooked due to experimental design or statistical
30 20 treatment of data. The study has also important
implications for quantitative genetics theory and
practical pig breeding.

35 *IGF2* was identified as a positional candidate gene
for this QTL due to the observed similarity between pig
25 chromosome 2p and human chromosome 11p. A genomic *IGF2*
clone was isolated by screening a porcine BAC library.
FISH analysis with this BAC clone gave a strong
40 consistent signal on the terminal part of chromosome 2p
(Fig. 1). A polymorphic microsatellite is located in the
30 3'UTR of *IGF2* in mice (GenBank U71085), humans (GenBank
S62623), and horse (GenBank AF020598). The possible
45 presence of a corresponding porcine microsatellite was
investigated by direct sequencing of the *IGF2* 3'UTR using
the BAC clone. A complex microsatellite was identified
50 35 about 800 bp downstream of the stop codon; a sequence
comparison revealed that this microsatellite is identical

5 to a previously described anonymous microsatellite,
Swc9⁶. PCR primers were designed and the microsatellite
(IGF2^{ms}) was found to be highly polymorphic with three
10 different alleles among the two Wild Boar founders and
5 another two among the eight Large White founders. IGF2^{ms}
was fully informative in the intercross as the breed of
origin as well as the parent of origin could be
15 determined with confidence for each allele in each F₂
animal.

10 A linkage analysis using the intercross pedigree was
carried out with IGF2^{ms} and the microsatellites Sw2443,
20 Sw2623, and Swr2516, all from the distal end of 2p⁷. IGF2
was firmly assigned to 2p by highly significant lod
scores (e.g. Z=89.0, θ =0.003 against Swr2516). Multipoint
15 analyses, including previously typed chromosome 2
25 markers⁸, revealed the following order of loci (sex-
average map distances in Kosambi cM): Sw2443/Swr2516-0.3-
IGF2-14.9-Sw2623-10.3-Sw256. No recombinant was observed
30 between Sw2443 and Swr2516, and the suggested proximal
20 location of IGF2 in relation to these loci is based on a
single recombinant giving a lod score support of 0.8 for
the reported order. The most distal marker in our
35 previous QTL study, Sw256, is located about 25 cM from
the distal end of the linkage group.

25 QTL analyses of body composition, fatness, meat
quality, and growth traits were carried out with the new
40 chromosome 2 map using a statistical model testing for
the possible presence of an imprinting effect as expected
for IGF2. Clear evidence for a paternally expressed QTL
30 located at the very distal tip of 2p was obtained (Fig.
45 2; Table 1). The QTL had very large effects on lean meat
content in ham and explained an astonishing 30% of the
residual phenotypic variance in the F₂ population. Large
50 effects on the area of the longissimus dorsi muscle, on
35 the weight of the heart, and on back-fat thickness

(subcutaneous fat) were also noted. A moderate effect on one meat quality trait, reflectance value, was indicated. The QTL had no significant effect on abdominal fat, birth weight, growth, weight of liver, kidney, or spleen (data not shown). The Large White allele at this QTL was associated with larger muscle mass and reduced back-fat thickness consistent with the difference between this breed and the Wild Boar population. The strong imprinting effect observed for all affected traits strongly suggests a single causative locus. The pleiotropic effects on skeletal muscle mass and the size of the heart appear adaptive from a physiological point of view as a larger muscle mass requires a larger cardiac output. The clear paternal expression of this QTL is illustrated by the least squares means which fall into two classes following the population origin of the paternally inherited allele (Table 1). It is worth noticing though that there was a non-significant trend towards less extreme values for the two heterozygous classes, in particular for the estimated effect on the area of longissimus dorsi. This may be due to chance, but could have a biological explanation, e.g. that there is some expression of the maternally inherited allele or that there is a linked, non-imprinted QTL with minor effects on the traits in question. ~

The *IGF2*-linked QTL and the *FAT1* QTL on chromosome 4 1, 9 are by far the two loci with the largest effect on body composition and fatness segregating in this Wild Boar intercross. The *IGF2* QTL controls primarily muscle mass whereas *FAT1* has major effects on fat deposition including abdominal fat, a trait that was not affected by the *IGF2* QTL (Fig. 2). No significant interaction between the two loci was indicated and they control a very large proportion of the residual phenotypic variance in the F_2 generation. A model including both QTLs explains 33.1% of the variance for percentage lean meat in ham, 31.3% for the percentage of lean meat plus bone in back, and 26.2%

5 for average back fat depth (compare with a model
including only chromosome 2 effects, Table 1). The two
QTLs must have played a major role in the response during
10 selection for lean growth and muscle mass in the Large
5 White domestic pig.

A highly significant segregation distortion was
observed in the *IGF2* region (excess of Wild Boar-derived
15 alleles) as shown in Table 1 ($\chi^2=11.7$, d.f.=2; $P=0.003$).
The frequency of Wild Boar-derived *IGF2* alleles was 59%
10 in contrast to the expected 50% and there was twice as
many "Wild Boar" as "Large White" homozygotes. This
20 deviation was observed with all three loci at the distal
tip and is thus not due to typing errors. The effect was
also observed with other loci but the degree of
15 distortion decreased as a function of the distance to the
distal tip of the chromosome. Blood samples for DNA
25 preparation were collected at 12 weeks of age and we are
convinced that the deviation from expected Mendelian
ratios was present at birth as the number of animals lost
30 prior to blood sampling was not sufficient to cause a
deviation of this magnitude. No other of the more than
250 loci analyzed in this pedigree show such a marked
35 segregation distortion (L. Andersson, unpublished). The
segregation distortion did not show an imprinting effect,
25 as the frequencies of the two reciprocal types of
heterozygotes were identical (Table 1). This does not
40 exclude the possibility that the QTL effects and the
segregation distortion are controlled by the same locus.
The segregation distortion maybe due to meiotic drive
30 favoring the paternally expressed allele during
gametogenesis, as the F_1 parents were all sired by Wild
45 Boar males. Another possibility is that the segregation
distortion may be due to codominant expression of the
maternal and paternal allele in some tissues and/or
50 during a critical period of embryo development. Biallelic
35 *IGF2* expression has been reported to occur to some extent

5 during human development^{10, 11} and interestingly a strong
influence of the parental species background on *IGF2*
expression was recently found in a cross between *Mus*
10 *musculus* and *Mus spretus*¹². It is also interesting that a
5 VNTR polymorphism at the insulin gene, which is very
closely linked to *IGF2*, is associated with size at birth
in humans¹³. It is possible that the *IGF2*-linked QTL in
15 pigs has a minor effect on birth weight but in our data
it was far from significant (Fig. 2) and there was no
10 indication of an imprinting effect.

This study is an advance in the general knowledge
20 concerning the biological importance of the *IGF2* locus.
The important role of *IGF2* for prenatal development is
well-documented from knock-out mice¹⁴ as well as from its
25 causative role in the human Beckwith-Wiedemann
syndrome¹⁵. This study demonstrates an important role for
the *IGF2*-region also for postnatal development. It should
be stressed that our intercross between outbred
30 populations is particularly powerful to detect QTL with a
parent of origin-specific effect on a multifactorial
20 trait. This is because multiple alleles (or haplotypes)
are segregating and we could deduce whether a
heterozygous *F₂* animal received the Wild Boar allele from
35 the *F₁* male or female. It is quite possible that the
segregation of a paternally expressed *IGF2*-linked QTL
25 affecting a trait like obesity has been overlooked in
human studies or in intercrosses between inbred rodent
40 populations because of experimental design or statistical
treatment of data. An imprinting effect cannot be
30 detected in an intercross between two inbred lines as
only two alleles are segregating at each locus. Our
45 result has therefore significant bearings on the future
analysis of the association between genetic polymorphism
50 in the *insulin-IGF2* region and Type I diabetes¹⁶,
35 obesity¹⁷, and variation in birth weight¹³ in humans, as

5 well as for the genetic dissection of complex traits
using inbred rodent models. A major impetus for
generating an intercross between the domestic pig and its
wild ancestor was to explore the possibilities to map and
10 5 identify major loci that have responded to selection. We
have now showed that two single QTLs on chromosome 2
(this study) and 4¹, 2 explain as much as one third of
the phenotypic variance for lean meat content in the F₂
15 generation. This is a gross deviation from the underlying
20 assumption in the classical infinitesimal model in
quantitative genetics theory namely that quantitative
traits are controlled by an infinite number of loci each
with an infinitesimal effect. If a large proportion of
the genetic difference between two divergent populations
25 (e.g. Wild Boar and Large White) is controlled by a few
loci, one would assume that selection would quickly fix
QTL alleles with large effects leading to a selection
plateau. However, this is not the experience in animal
breeding programs or selection experiments where good
30 20 persistent long-term selection responses are generally
obtained, provided that the effective population size is
reasonably large¹⁸. A possible explanation for this
paradox is that QTL alleles controlling a large
35 proportion of genetic differences between two populations
25 may be due to several consecutive mutations; this may be
mutations in the same gene or at several closely linked
genes affecting the same trait. It has been argued that
40 new mutations contribute substantially to long-term
selection responses¹⁹, but the genomic distribution of
30 such mutations are unknown.

45 The search for a single causative mutation is the
paradigm as regards the analysis of genetic defects in
mice and monogenic disorders in humans. We propose that
this may not be the case for loci that have been under
50 35 selection for a large number of generations in domestic
animals, crops, or natural populations. This hypothesis

5 predicts the presence of multiple alleles at major QTL.
It gains some support from our recent characterization of
porcine coat color variation. We have found that both the
alleles for dominant white color and for black-spotting
10 differ from the corresponding wild-type alleles by at
least two consecutive mutations with phenotypic effects
at the *KIT* and *MC1R* loci, respectively^{20, 21}. In this
context it is highly interesting that in the accompanying
example we have identified a third allele at the *IGF2*-
15 linked QTL. The effects on muscle mass of the three
alleles rank in the same order as the breeds in which
they are found i.e. Piétrain pigs are more muscular than
20 Large White pigs that in turn have higher lean meat
content than Wild Boars.

15 There are good reasons to decide that *IGF2* is the
causative gene for the now reported QTL. Firstly, there
is a perfect agreement in map localization (Fig. 2).
Secondly, it has been shown that *IGF2* is paternally
expressed in mice, humans, and now in pigs, like the QTL.

20 There are several other imprinted genes in the near
vicinity of *IGF2* in mice and humans (*Mash2*, *INS2*, *H19*,
KVLQT1, *TAPA1/CD81*, and *CDKN1C/p57^{KIP2}*) but only *IGF2* is
35 paternally expressed in adult tissues²². We believe that
this locus provides a unique opportunity for molecular
25 characterization of a QTL. The clear paternal expression
can be used to exclude genes that do not show this mode
of inheritance. Moreover, the presence of an allelic
40 series should facilitate the difficult distinction
between causative mutations and linked neutral
30 polymorphism. We have already shown that there is no
45 difference in coding sequence between *IGF2* alleles from
Piétrain and Large White pigs suggesting that the
causative mutations occur in regulatory sequences. An
obvious step is to sequence the entire *IGF2* gene and its
50 35 multiple promoters from the three populations. The recent

5 report that a VNTR polymorphism in the promoter region of
the insulin (*INS*) gene affects *IGF2* expression²³ suggests
that the causative mutations may be at a considerable
10 distance from the *IGF2* coding sequence.

5 The results have several important implications for
the pig breeding industry. They show that genetic
imprinting is not an esoteric academic question but need
to be considered in practical breeding programs. The
15 detection of three different alleles in Wild Boar, Large
20 White, and Piétrain populations indicates that further
alleles at the *IGF2*-linked QTL segregate within
commercial populations. The paternal expression of the
QTL facilitates its detection using large paternal half-
sib families as the female contribution can be ignored.
15 The QTL is exploited to improve lean meat content by
25 marker assisted selection within populations or by marker -
assisted introgression of favorable alleles from one
population to another.

Example 2: Piétrain x Large White intercrosses

Methods

Pedigree material: The pedigree material utilized to map

QTL was selected from a previously described Piétrain x Large White F2 pedigree comprising > 1,800 individuals^{6,7}.

To assemble this F2 material, 27 Piétrain boars were mated to 20 Large White sows to generate an F1 generation comprising 456 individuals. 31 F1 boars were mated to

unrelated 82 F1 sows from 1984 to 1989, yielding a total of 1862 F2 offspring. F1 boars were mated on average to 7 females, and F1 sows to an average of 2,7 males. Average offspring per boar were 60 and per sow 23.

Phenotypic information: (i) *Data collection:* A total of 21 distinct phenotypes were recorded in the F2 generation^{6,7}. These included:

- five growth traits: birth weight (g), weaning weight (Kg), grower weight (Kg), finisher weight (Kg) and average daily gain (ADG; Kg/day; grower to finisher period);

- two body proportion measurements: carcass length (cm); and a conformation score (0 to 10 scale; ref.6);

- ten measurements of carcass composition obtained by dissection of the chilled carcasses 24 hours after slaughter. These include measurements of muscularity: % ham (weight hams/carcass weight), % loin (weight loin/carcass weight), % shoulder (weight shoulder/carcass weight), % lean cuts (% ham + %loin + % shoulder); and measurements of fatness: average back-fat thickness (BFT; cm), % backfat (weight backfat/carcass weight), % belly (weight belly/carcass weight), % leaf fat (weight leaf fat/carcass weight), % jowl (weight jowl/carcass weight), and "% fat cuts" (% backfat + % belly + % leaf fat + % jowl).

- four meat quality measurements: pH _{LM1} (*Longissimus dorsi* 1

hour after slaughter), pH_{LD24} (*Longissimus dorsi* 24 hours after slaughter), pH_{G1} (*Gracilis* 1 hour after slaughter) and pH_{G24} (*Gracilis* 24 hours after slaughter). (ii) *Data processing*: Individual phenotypes were preadjusted for fixed effects (sire, dam, CRC genotype, sex, year-season, parity) and covariates (litter size, birth weight, weaning weight, grower weight, finisher weight) that proved to significantly affect the corresponding trait. Variables included in the model were selected by stepwise regression.

Marker genotyping: Primer pairs utilized for PCR amplification of microsatellite markers are as described¹⁹. Marker genotyping was performed as previously described²⁰. Genotypes at the *CRC* and *MyoD* loci were determined using conventional methods as described^{1,12}. The LAR test for the *Igf2* SNP was developed according to Baron et al.²¹ using a primer pair for PCR amplification (5'-CCCTGAACTTGAGGACGAGCAGCC-3'; 5'-ATCGCTGTGGGCTGGGTGGGCTGCC-3') and a set of three primers for the LAR step (5'-FAM-CGCCCCAGCTGCCCCCAG-3'; 5'-HEX-CGCCCCAGCTGCCCCCAA-3'; 5'-CCTGAGCTGCAGCAGGCCAG-3').

Map construction: Marker maps were constructed using the TWOPOINT, BUILD and CHROMPIC options of the CRIMAP package²². To allow utilisation of this package, full-sib families related via the boar or sow were disconnected and treated independently. By doing so, some potentially usable information was neglected, yielding, however, unbiased estimates of recombination rates.

QTL mapping: (i) *Mapping Mendelian QTL*: Conventional QTL mapping was performed using a multipoint maximum likelihood method. The applied model assumed one segregating QTL per

chromosome, and fixation of alternate QTL alleles in the respective parental lines, Piétrain (P) and Large White (LW). A specific analysis program had to be developed to account for the missing genotypes of the parental generation, resulting in the fact that the parental origin of the F1 chromosomes could not be determined. Using a typical "interval mapping" strategy, an hypothetical QTL was moved along the marker map using user-defined steps. At each position, the likelihood (L) of the pedigree data was computed as:

$$L = \sum_{\theta=1}^{2^r} \prod_{i=1}^n \sum_{G=1}^4 (P(G|M_i, \theta, \phi) P(y_i|G))$$

P or right chromosome P), there is a total of 2^r combinations for r F1 parents.

$$\prod_{i=1}^n n \text{ F2}$$

$\sum_{G=1}^4$ i th F2 offspring, over the four possible QTL genotypes:

P/P , P/LW , LW/P and LW/LW

$P(G|M_i, \theta, \phi)$: the marker genotype of the i th F2 offspring and its F1 parents, (ii) : the vector of recombination rates between adjacent markers and between the hypothetical QTL and its flanking markers, and (iii) θ the considered marker-QTL phase combination of the F1 parents.

Recombination rates and marker linkage phase of F1 parents are assumed to be known when computing this probability. Both were determined using CRIMAP in the map construction phase (see above).

$P(y_i|G)$ of offspring i , given the QTL genotype under consideration. This probability is computed from the normal density function:

$$P(y_i|G) = \frac{1}{\sqrt{2\pi}\sigma} e^{-\frac{(y_i - \mu_G)^2}{2\sigma^2}}$$

μ_G is the phenotypic mean of the considered QTL genotype (PP, PL, LP or LL) and σ^2 the residual variance σ^2 was considered to be the same for the four QTL genotypic classes.

The values of μ_{PP} , $\mu_{PL}=\mu_{LP}$, μ_{LL} and σ^2 maximizing L were determined using the GEMINI optimisation routine²³.

The likelihood obtained under this alternative H_1 hypothesis was compared with the likelihood obtained under the null hypothesis H_0 of no QTL, in which the phenotypic means of the four QTL genotypic classes were forced to be identical. The difference between the logarithms of the corresponding likelihoods yields a lodscore measuring the evidence in favour of a QTL at the corresponding map position.

(ii) *Significance thresholds*: Following Lander & Botstein²⁴, lodscore thresholds (T) associated with a chosen genome-wise significance level, were computed such that:

$$\alpha = (C + 9.21GT) \chi^2_1(4.6T)$$

C corresponds to the number of chromosomes (= 19), G corresponds to the length of the genome in Morgans (= 29),

and $\chi^2_1(4.6T)$ denotes one minus the cumulative distribution function of the chi-squared distribution with 2 d.f. Single point $2\ln(LR)$ were assumed to be distributed as a chi-squared distribution with two degrees of freedom, as we were fitting both an additive and dominance component. To account for the fact that we were analysing multiple traits, significance levels were adjusted by applying a Bonferoni correction corresponding to the effective number of independent traits that were analyzed. This effective number was estimated at 16 following the approach described by Speiman et al.²⁵.

Altogether, this allowed us to set the lodscore threshold associated with an experiment-wise significance level of 5%

at 5.8. When attempting to confirm the identified QTL in an independent sample, the same approach was used, however, setting C at 1, G at 25cM and correcting for the analysis of 4.5 independent traits (as only six traits were analyzed in this sample). This yielded a lodscore threshold associated with a Type I error of 5% of 2.

(iii). *Testing for an imprinted QTL:* To test for an imprinted QTL, we assumed that only the QTL alleles transmitted by the parent of a given sex would have an effect on phenotype, the QTL alleles transmitted by the other parent being "neutral". The likelihood of the pedigree data under this hypothesis was computed using equation 1. To compute $P(y_i | G)$, however, the phenotypic means of the four QTL genotypes were set at $\mu_{PP} = \mu_{PL} = \mu_P$ and $\mu_{LP} = \mu_{LL} = \mu_L$ to test for a QTL for which the paternal allele only is expressed, and $\mu_{PP} = \mu_{LP} = \mu_P$ and $\mu_{PL} = \mu_{LL} = \mu_L$ to test for a QTL for which the maternal allele only is expressed. It is assumed in this notation that the first subscript refers to the paternal allele, the second subscript to the maternal allele. H_0 was defined as the null-hypothesis of no QTL, H_1 testing the presence of a Mendelian QTL; H_2 testing the presence of a paternally expressed QTL, and H_3 testing the presence of a maternally expressed QTL.

RT-PCR: Total RNA was extracted from skeletal muscle according to Chirgwin et al.²⁶. RT-PCR was performed using the Gene-Amp RNA PCR Kit (Perkin-Elmer). The PCR products were purified using QiaQuick PCR Purification kit (Qiagen) and sequenced using Dye terminator Cycle Sequencing Ready Reaction (Perkin Elmer) and an ABI373 automatic sequencer.

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In example 2 we report the identification of a QTL with major effect on muscle mass and fat deposition mapping to porcine 2p1.7. The QTL shows clear evidence for parental imprinting strongly suggesting the involvement of the *Igf2* locus.

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5 A Piétrain X Large White intercross comprising 1125 F_2 offspring was generated as described^{6,7}. The Large White and Piétrain parental breeds differ for a number of economically important phenotypes. Piétrains are famed for their exceptional muscularity and leanness⁸ (Figure 2), while Large
10 Whites show superior growth performance. Twenty-one distinct phenotypes measuring (i) growth performance (5), (ii) muscularity (6), (iii) fat deposition (6), and (iv) meat quality (4), were recorded on all F_2 offspring.

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In order to map QTL underlying the genetic differences
15 between these breeds, we undertook a whole genome scan using microsatellite markers on an initial sample of 677 F_2 individuals. Analysis of pig chromosome 2 using a ML multipoint algorithm, revealed highly significant lodscores (up to 20) for six of the 12 phenotypes measuring muscularity
30 and fat deposition at the distal end of the short arm of chromosome 2 (Figure 3a). Positive lodscores were obtained for the remaining six phenotypes, however, not reaching the genome-wide significance threshold ($\alpha = 5\%$). To confirm this finding, the remaining sample of 355 F_2 offspring was
25 genotyped for the five most distal 2p markers and QTL analysis performed for the traits yielding the highest lodscores in the first analysis. Lodscores ranged from 2.1 to 7.7, clearly confirming the presence of a major QTL in this region. Table 2 reports the corresponding ML estimates for
40 the three genotypic means as well as the corresponding residual variance.

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30 Bidirectional chromosome painting establishes a correspondence between SSC2p and HSA11pter-q13^{9,10}. At least

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two serious candidate genes map to this region in man: the myogenic basic helix-loop-helix factor, *MyoD*, maps to HSA11p15.4, while *Igf2* maps to HSA11p15.5. *MyoD* is a well known key regulator of myogenesis and is one of the first myogenic markers to be switched on during development¹¹. A previously described amplified sequence polymorphism in the porcine *MyoD* gene¹² proved to segregate in our F₂ material, which was entirely genotyped for this marker. Linkage analysis positioned the *MyoD* gene in the SW240-SW776 (odds > 1000) interval, therefore well outside the lod-2 drop off support interval for the QTL (figure 1). *Igf2* is known to enhance both proliferation and differentiation of myoblasts *in vitro*¹³ and to cause a muscular hypertrophy when overexpressed *in vivo*. Based on a published porcine adult liver cDNA sequence¹⁴, we designed primer pairs allowing us to amplify the entire *Igf2* coding sequence with 222 bp of leader and 280 bp of trailer sequence from adult skeletal muscle cDNA. Piétrain and Large White RT-PCR products were sequenced indicating that the coding sequences was identical in both breeds and with the published sequence. However, a G A transition was found in the leader sequence corresponding to exon 2 in man (Figure 4). We developed a screening test for this single nucleotide polymorphism (SNP) based on the ligation amplification reaction (LAR), allowing us to genotype our pedigree material. Based on these data, *Igf2* was shown to colocalize with the SWC9 microsatellite marker (= 0%), therefore located at approximately 2 centimorgan from the most likely position of the QTL and well within the 95% support interval for the QTL (figure 1). Subsequent sequence analysis demonstrated that the microsatellite marker SWC9 is actually located within the 3' UTR of the *Igf2* gene. Combined with available comparative mapping data for the PGA and FSH loci, these results suggest the occurrence of an interstitial

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inversion of a chromosome segment containing *MyoD*, but not *Igf2* which has remained telomeric in both species.

Igf2 therefore appeared as a strong positional allele having the observed QTL effect. In man and mouse, *Igf2* is known to be imprinted and to be expressed exclusively from the paternal allele in several tissues¹⁵. Analysis of skeletal muscle cDNA from pigs heterozygous for the SNP and/or SWC9, shows that the same imprinting holds in this tissue in the pig as well (Figure 4). Therefore if *Igf2* were responsible for the observed effect, and knowing that only the paternal *Igf2* allele is expressed, one can predict that (i) the paternal allele transmitted by F1 boars (P or LW) would have an effect on phenotype of F2 offspring, (ii) the maternal allele transmitted by F1 sows (P or LW) would have no effect on phenotype of F2 offspring, and (iii) the likelihood of the data would be superior under a model of a bimodal (1:1) F2 population sorted by inherited paternal allele when compared to a conventional "Mendelian" model of a trimodal (1:2:1) F2 population. The QTL mapping programs were adapted in order to allow testing of the corresponding hypotheses. H_0 was defined as the null-hypothesis of no QTL, H_1 as testing for the presence of a Mendelian QTL, H_2 as testing for the presence of a paternally expressed QTL, and H_3 as testing for the presence of a maternally expressed QTL.

Figure 3 summarizes the obtained results. Figure 3a, 3b and 3c respectively show the lodscore curves corresponding to $\log_{10} (H_2/H_0)$, $\log_{10} (H_3/H_0)$ and $\log_{10} (H_2/H_1)$. It can be seen that very significant lodscores are obtained when testing for the presence of a paternally expressed QTL, while there is no evidence at all for the segregation of a QTL when studying the chromosomes transmitted by the sows. Also, the hypothesis of a paternally expressed QTL is significantly more likely ($\log_{10} (H_2/H_1) > 3$) than the hypothesis of a "Mendelian" QTL

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for all examined traits. The fact that the same tendency is observed for all traits indicates that it is likely the same imprinted gene that is responsible for the effects observed on the different traits. Table 2 reports the ML phenotypic means for the F2 offspring sorted by inherited paternal QTL allele. Note that when performing the analysis under a model of a mendelian QTL, the Piétrain and Large White QTL alleles appeared to behave in an additive fashion, the heterozygous genotype exhibiting a phenotypic mean corresponding exactly to the midpoint between the two homzygous genotypes. This is exactly what one would predict when dealing with an imprinted QTL as halve of the heterozygous offspring are expected to have inherited the P allele from their sire, the other halve the LW allele.

These data therefore confirmed our hypothesis of the involvement of an imprinted gene expressed exclusively from the paternal allele. The fact that the identified chromosomal segment coincides precisely with an imprinted domain documented in man and mice strongly implicates the orthologous region in pigs. At least seven imprinted genes mapping to this domain have been documented (*Igf2*, *Ins2*, *H19*, *Mash2*, *p57^{KIP2}*, *KvLQTL1* and *TDAG51*) (ref. 15 and Andrew Feinberg, personal communication). Amongst these, only *Igf2* and *Ins2* are paternally expressed. While we cannot exclude that the observed QTL effect is due to an as of yet unidentified imprinted gene in this region, its reported effects on myogenesis *in vitro* and *in vivo*¹³ strongly implicate *Igf2*. Particularly the muscular hypertrophy observed in transgenic mice overexpressing *Igf2* from a muscle specific promotor are in support of this hypothesis (Nadia Rosenthal, personal communication. Note that allelic variants of the *INS* VNTR have recently been shown to be associated

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with size at birth¹⁶ in man¹⁶, and that the same VNTR has been shown to affect the level of *Igf2* expression¹⁷.

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The observation of the same QTL effect in a Large White x Wild Boar intercross indicates the existence of a series of at least three distinct functional alleles. Moreover, preliminary evidence based on marker assisted segregation analysis points towards residual segregation at this locus within the Piétrain population (data not shown). The occurrence of an allelic series might be invaluable in identifying the causal polymorphisms which - based on the quantitative nature of the observed effect - are unlikely to be gross gene alterations but rather subtle regulatory mutations.

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The effects of the identified QTL on muscle mass and fat deposition are truly major, being of the same magnitude of those reported for the *CRC* locus^{6,7} though apparently without the associated deleterious effects on meat quality. We estimate that both loci jointly explain close to 50% of the Piétrain versus Large White breed difference for muscularity and leanness. Understanding the parent-of-origin effect characterizing this locus will allow for its optimal use in breeding programs. Indeed, today half of the offspring from commercially popular Piétrain x Large White crossbred boars inherit the unfavourable Large White allele causing considerable loss.

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The QTL described in this work is the second example of a gene affecting muscle development in livestock species that exhibits a non-mendelian inheritance pattern. Indeed, we have previously shown that the callipyge locus (related to the qualitative trait wherein muscles are doubled) is characterized by polar overdominance in which only the heterozygous individuals that inherit the CLPG mutation from their sire express the double-muscling phenotype⁵. This

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demonstrates that parent-of-origin effects affecting genes underlying production traits in livestock might be relatively common.

5 Example 3:

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Generating a reference sequence of IGF2 and flanking loci in the pig.

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10 The invention provides an imprinted QTL with major effect on muscle mass mapping to the IGF2 locus in the pig, and use of the QTL as tool in marker assisted selection. To fine tune this tool for marker assisted selection, as well as to further identify a causal mutation, we have further generated a reference sequence encompassing the entire porcine IGF2 sequence as well as that from flanking genes.

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To achieve this, we screened a porcine BAC library with IGF2 probes and identified two BACs. BAC-PIGF2-1 proved to

20 contain the INS and IGF2 genes, while BAC-PIGF2-2 proved to contain the IGF2 and H19 genes. The NotI map as well as the relative position of the two BACs is shown in Figure 5. BAC-PIGF2-1 was shotgun sequenced using standard procedures and automatic sequencers. The resulting sequences were assembled using standard software yielding a total of 115 contigs. The corresponding sequences are reported in figure 6. Similarity searches were performed between the porcine contigs and the orthologous sequences in human. Significant homologies were detected for 18 contigs and are reported in Figure 7.

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For BAC-PIGF2-2, the 24 Kb NotI fragment not present in BAC-PIGF2-1 was subcloned and sequenced using the EZ::TN transposon approach and ABI automatic sequencers. Resulting

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sequences were assembled using the Phred-Phrap-Consed program
suit, yielding seven distinct contigs (figure 8). The contig
sequences were aligned with the corresponding orthologous
human sequences using the compare and dotplot programs of the
5 GCG suite. Figure 9 summarizes the corresponding results.

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Example 4: Identification of DNA sequence polymorphisms in
the IGF2 and flanking loci.

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10 Based on the reference sequence obtained as described in
Example 1, we resequenced part of the IGF2 and flanking loci
from genomic DNA isolated from Piétrain, Large White and Wild
Boar individuals, allowing identification of DNA sequence
polymorphisms such as reported in figure 10.

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Legends to the figures

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Fig. 1: Test statistic curves obtained in QTL analyses of chromosome 2 in a Wild Boar/Large White intercross. The graph plots the F ratio testing the hypothesis of a single QTL at a given position along the chromosome for the traits indicated. The marker map with the distances between markers in Kosambi centiMorgan is given on the X-axis. The horizontal lines represent genome-wise significant ($P < 0.05$) and suggestive levels for the trait lean meat in ham; similar significance thresholds were obtained for the other traits.

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Figure 2: Piétrain pig with characteristic muscular hypertrophy.

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Figure 3: Lodscore curves obtained in a Piétrain x Large White intercross for six phenotypes measuring muscle mass and fat deposition on pig chromosome 2. The most likely positions of the *Igf2* and *MyoD* genes determined by linkage analysis with respect to the microsatellite marker map are shown. H_0 was defined as the null-hypothesis of no QTL, H_1 as testing for the presence of a Mendelian QTL, H_2 as testing for the presence of a paternally expressed QTL, and H_3 as testing for the presence of a maternally expressed QTL. 3a: $\log_{10}(H_1/H_0)$, 3b: $\log_{10}(H_2/H_0)$, 3c: $\log_{10}(H_3/H_0)$

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Figure 4: A. Structure of the human *Igf2* gene according to ref. 17, with aligned porcine adult liver cDNA sequence as reported in ref. 16. The position of the nt241(G-A) transition and *Swc9* microsatellite are shown. B. The corresponding markers were used to demonstrate the monoallelic (paternal) expression of *Igf2* in skeletal muscle

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and liver of 10-week old fetuses. PCR amplification of the *nt421*(G-A) polymorphism and *Swc9* microsatellite from genomic DNA clearly shows the heterozygosity of the fetus, while only the paternal allele is detected in liver cDNA (*nt421*(G-A) and *Swc9*) and muscle cDNA (*Swc9*). The absence of RT-PCR product for *nt421*(G-A) from in fetal muscle points towards the absence of mRNA including exon 2 in this tissue. Parental origin of the foetal alleles was determined from the genotypes of sire and dam (data not shown).

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Figure 5: A NotI restriction map showing the relative position of BAC-PIGF2-1 (comprising INS and IGF2 genes), and BAC-PIGF2-2 (comprising IGF2 and H19 genes).

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15 Figure 6: Nucleic acid sequences of contig 1 to contig 115 derived from BAC-PIGF2-1 which was shotgun sequenced using standard procedures and automatic sequencers.

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Figure 7: Similarity between porcine contigs of figure 6 and orthologous sequences in human.

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Figure 8 Nucleic acid sequences of contig 1 to contig 7 derived from BAC-PIGF2-2, (the 24 Kb NotI fragment not present in BAC-PIGF2-1) which was subcloned and sequenced using the E2::TN transposon approach and ABI automatic sequencers.

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Figure 9: Similarity between porcine contigs of figure 8 and orthologous sequences in human.

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Figure 10: DNA sequence polymorphisms in the IGF2 and flanking loci from genomic DNA isolated from Pietrain, Large White and Wild Boar individuals.

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REFERENCES

Literature cited with example 1

1. Andersson, L. et al. Genetic mapping of quantitative trait loci for growth and fatness in pigs. *Science* 263, 1771-1774 (1994).
2. Knott, S.A. et al. Multiple marker mapping of quantitative trait loci in a cross between outbred wild boar and Large White pigs. *Genetics* 149, 1069-1080 (1998).
3. Edfors-Lilja, I. et al. Mapping quantitative trait loci for immune capacity in the pig. *Journal of Immunology* 161, 829-835 (1998).
4. Andersson-Eklund, L. et al. Mapping quantitative trait loci for carcass and meat quality traits in a wild boar x Large White intercross. *Journal of Animal Science* 76, 694-700 (1998).
5. Fronicke, L., Chowdhary, B.P., Scherthan, H. & Gustavsson, I. A comparative map of the porcine and human genomes demonstrates ZOO-FISH and gene mapping-based chromosomal homologies. *Mamm Genome* 7, 285-90 (1996).
6. Alexander, L.J. et al. Physical assignments of 68 porcine cosmids and lambda clones containing microsatellites. *Mammalian Genome* 7, 368-372 (1996).
7. Rohrer, G.A. et al. A comprehensive map of the porcine genome. *Genome Research* 6, 371-391 (1996).
8. Marklund, L. et al. A comprehensive linkage map of the pig based on a wild pig-Large White intercross. *Anim Genet* 27, 255-69 (1996).
9. Marklund, L., Nyström, P.E., Stern, S., Andersson-Eklund, L. & Andersson, L. Quantitative trait loci for

fatness and growth on pig chromosome 4. *Heredity* In press(1998).

10. Ohlsson, R., Hedborg, F., Holmgren, L., Walsh, C. & Ekstrom, T.J. Overlapping patterns of IGF2 and H19 expression during human development: biallelic IGF2 expression correlates with a lack of H19 expression. *Development* 120, 361-368 (1994).
11. Ekström, T.J., Cui, H., Li, X. & Ohlsson, R. Promoter-specific IGF2 imprinting status and its plasticity during human liver development. *Development* 121, 309-316 (1995).
12. Hemberger, M. et al. H19 and Igf2 are expressed and differentially imprinted in neuroectoderm-derived cells in the mouse brain. *Dev. Genes Evol.* 208, 393-402 (1998).
13. Dunger, D.B. et al. Association of the INS VNTR with size at birth. *Nature Genetics* 19, 98-100 (1998).
14. DeChiara, T.M., Robertson, E.J. & Efstratiadis, A. Parental imprinting of the mouse insulin-like growth factor II gene. *Cell* 64, 849-859 (1991).
15. Sun, F.L., Dean, W.L., Kelsey, G., Allen, N.D. & Reik, W. Transactivation of Igf2 in a mouse model of Beckwith-Wiedemann syndrome. *Nature* 389, 809-815 (1997).
16. Davies, J.L. et al. A genome-wide search for human type 1 diabetes susceptibility genes. *Nature* 371, 130-136 (1994).
17. O'Dell, S.D. et al. ApaI polymorphism in insulin-like growth factor II (IGF2) gene and weight in middle-aged males. *International Journal of Obesity* 21, 822-825 (1997).
18. Falconer, D.S. & Mackay, T.F.C. *Introduction to Quantitative Genetics*, (Longman, England, 1996).
19. Hill, W.G. Rates of change in quantitative traits from fixation of new mutations. *Proc Natl Acad Sci U S A* 79, 142-145 (1982).

20. Marklund, S. et al. Molecular basis for the dominant white phenotype in the domestic pig. *Genome Research* 8, 826-833 (1998).
21. Kijas, J.M.H. et al. Melanocortin receptor 1 (MC1R) mutations and coat color in the pig. *Genetics* In press(1998).
22. Beechey, C.V. personal communication (1998).
23. Paquette, J., Giannoukakis, N., Polychronakos, C., Vafiadis, P. & Deal, C. The *INS* 5' variable number of tandem repeats is associated with *IGF2* expression in humans. *Journal of Biological Chemistry* 273, 14158-14164 (1998).
24. Sambrook, J., Fritsch, E.F. & Maniatis, T. *Molecular cloning : A laboratory manual.*, (Cold Spring Harbor Laboratory Press, Cold Spring Harbor, 1989).
25. Chowdhary, B.P., de la Sena, C., Harbitz, I., Eriksson, L. & Gustavsson, I. FISH on metaphase and interphase chromosomes demonstrates the physical order of the genes for GPI, CRC, and LIPE in pigs. *Cytogenetics Cell Genetics* 71, 175-178 (1995).
26. Green, P., Falls, K. & Crook, S. *Documentation for CRI-MAP, version 2.4.*, (Washington University School of Medicine, St Louise, MO, 1990).
27. Haley, C.S., Knott, S.A. & Elsen, J.M. Mapping quantitative trait loci in crosses between outbred lines using least squares. *Genetics* 136, 1195-1207 (1994).
28. Churchill, G.A. & Doerge, R.W. Empirical threshold values for quantitative trait mapping. *Genetics* 138, 963-971 (1994).
29. Anonymous. *SAS version 6.10*, (SAS Institute Inc., Cary, NC., 1990).

References used with example 2:

1. Fuji, J.; Otsu, K.; Zorzato, F.; Deleon, S.; Khanna, V.K.; Weiler, J.E. O'Brien, P.J.; MacLennan, D.H. (1991). Identification of a mutation in the porcine ryanodine receptor associated with malignant hyperthermia. *Science* 253: 448-451.
2. MacLennan, D.H. & Phillips, M.S. (1993). Malignant hyperthermia. *Science* 256:789-794.
3. Grobet, L.; Royo Martin, L.J.; Poncelet, D.; Pirottin, D.; Brouwers, B.; Riquet, J.; Schoeberlein, A.; Dunner, S.; Ménissier, F.; Massabanda, J.; Fries, R.; Hanset, R.; Georges, M. (1997). A deletion in the myostatin gene causes double-muscling in cattle. *Nature Genetics* 17:71-74.
4. Andersson, L.; Haley, C.S.; Ellegren, H.; Knott, S.A.; Johansson, M.; Andersson, K.; Andersson-Eklund, L.; Edfors-Lilja, I.; Fredholm, M.; Hansson, I.; Håkansson, J.; Lundström, K. (1994). Genetic mapping of quantitative trait loci for growth and fatness in pigs. *Science* 263:1771-1774.
5. Cockett, N.; Jackson, S.; Shaw, T.; Farnir, F.; Berghmans, S.; Snowden, G.; Nielsen, D.; Georges, M. (1996). Polar overdominance at the ovine callipyge locus. *Science* 273:236-238.
6. Hanset, R.; Dasnois, C.; Scalais, S.; Michaux, C.; Grobet, L. (1995). Genotypes at the locus for halothane sensitivity and performance in a Piétrain x Large White F2. *Genet. Sel. Evol.* 27: 63-76.
7. Hanset, R.; Dasnois, C.; Scalais, S.; Michaux, C.; Grobet, L. (1995). Introgression into the Piétrain genome of the normal allele at the locus for halothane sensitivity. *Genet. Sel. Evol.* 27: 77-88.
8. Olivier, L.; Lauvergne, J.J. (1967). A study of the inheritance of the muscular hypertrophy of the Piétrain pig: preliminary results. *Annales de Médecine Vétérinaire* 111: 104-109.

5

50

10

9. Rettenberger, G.; Klett, C.; Zechner, U.; Kunz, J.; Vogel, W.; Hameister, H. (1995). Visualisation of the conservation of synteny between humans and pigs by heterologous chromosome painting. *Genomics* 26: 372-378.

15

5 10. Goureau, A.; Yerle, M.; Schmitz, A.; Riquet, J.; Milan, D.; Pinton, P.; Frelat, G.; Gellin, J. (1996). Human and porcine correspondence of chromosome segments using bidirectional chromosome painting. *Genomics* 36:252-262.

20

11. Yun, K.; Wold, B. (1996). Skeletal muscle determination and differentiation: story of a core regulatory network and its context. *Current Opinion in Cell Biology* 8:877-889.

25

12. Knoll, A.; Nebola, M.; Dvorak, J.; Cepica, S. (1997). Detection of a DdeI PCR RFLP within intron 1 of the porcine MYOD1(MYF3) locus. *Animal Genetics* 28, 308-322.

30

13. Florini, J.R.; Ewton, D.Z.; McWade, F.J. (1995). IGFs, muscle growth, and myogenesis. *Diabetes Review* 3:73-92.

35

14. Catchpole, I.R.; Engström, W. (1990). Nucleotide sequence of a porcine insulin-like growth factor II cDNA. *Nucleic Acids Research* 18(21):6430.

40

15. Feil, R.; Moore, T.F.; Oswald, J.; Walter, J.; Sun, F.; Reik, W. (1997). The imprinted insulin like growth factor 2 gene. Pp70 In *Genomic Imprinting*. Eds. Reik & Surani. IRL Press at Oxford University Press.

45

16. Dunger, D.B.; Ong, K.K.L.; Huxtable, S.J.; Sherriff, A.; Woods, K.A.; Ahmed, M.L.; Golding, J.; Pembrey, M.E.; Ring, S.; the ALSPAC study team, Bennett, S.T.; Todd, J.A. (1998). Association of the INS VNTR with size at birth. *Nature Genetics* 19: 98-100.

50

17. Paquette J, Giannoukakis N, Polychronakos C, Vafiadis P, Deal C. (1998) The INS 5' variable number of tandem repeats is associated with IGF2 expression in humans. *J. Biol Chem* 273(23):14158-14164

55

18. Andersson-Eklund, L.; Marklund, L.; Lundström, K.; Haley, C.S.; Andersson, K.; Hansson, I.; Moller, M.; Andersson, L. (1998). Mapping Quantitative Trait Loci for carcass and meat quality traits in a Wild Boar x Large White intercross. *J. Anim. Sci.* 76:694-700.
19. Rohrer, G.A.; Alexander, L.J.; Hu, Z.; Keele, J.W.; Smith, T.P.; Beattie, C.W. (1996). A comprehensive map of the porcine genome. *Genome Research*, in the press.
20. Georges, M.; Nielsen, D.; Mackinnon, M.; Mishra, A.; Okimoto, R.; Pasquino, A.T.; Sargeant, L.S.; Sorensen, A.; Steele, M.R.; Zhao, X.; Womack, J.E.; Hoeschele, I. (1995). Mapping quantitative trait loci controlling milk production by exploiting progeny testing. *Genetics* 139: 907-920.
21. Baron, H.; Fung, S.; Aydin, A.; Bahring, S.; Luft, F.C.; Schuster, H. (1996). Oligonucleotide ligation assay (OLA) for the diagnosis of familial hypercholesterolemia. *Nat. Biotechnol.* 14(10):1279-1282.
22. Lander, E.; Green, P. (1987) Construction of multilocus genetic linkage maps in humans. *Proceedings of National Academy of Science (USA)* 84: 2363-2367.
23. Lalouel, J.M. (1983). Optimization of functions. *Contrib. Epidemiol.Biostat.* 4:235-259.
24. Lander, E.S. & Botstein, D. (1989). Mapping mendelian factors underlying quantitative traits using RFLP linkage maps. *Genetics* 121:185-199.
25. Spelman RL, Coppieters W, Karim L, van Arendonk JAM, Bovenhuis H (1996) Quantitative trait loci analysis for five milk production traits on chromosome six in the dutch Holstein-Friesian population. *Genetics* 144:1799-1808.
26. Chirgwin, J.M.; Przybyla, A.E.; MacDonald, R.J.; Rutter, W.J. (1979) Isolation of biologically active ribonucleic acid from sources enriched in ribonuclease. *Biochemistry* 18:5294-5299

Table 1 Summary of QTL analysis for pig chromosome 2 in a Wild Boar/Large White intercross

Trait	F ratio ²	QTL	Imprinting	Map position ³	Percent of F ₁ variance ⁴	Least squares means ¹		
						LP/LM	LP/WM	LP/MM
						n=62	n=43	n=43
						n=30		
<u>Body composition traits</u>								
Lean meat in ham, %	24.4***	19.1***	0	30.6	63.6 ^a	64.2 ^a	66.4 ^b	67.3 ^b
Lean meat mass in ham, kg	18.1***	16.8***	1	24.3	4.69 ^a	4.72 ^a	4.94 ^b	5.02 ^b
Lean meat + bone in back, %	12.2**	9.6**	0	17.4	66.3 ^a	66.7 ^a	69.3 ^b	70.8 ^b
Longissimus muscle area, cm ²	10.3**	4.8*	1	15.4	31.9 ^a	33.0 ^a	34.5 ^b	35.2 ^b
<u>Fatness traits</u>								
Average back fat depth, mm	7.1*	8.7**	0	10.4	27.2 ^a	27.7 ^a	25.5 ^b	24.7 ^b
<u>Weight of internal organs</u>								
Heart, gram	9.7**	11.4***	0	14.4	226 ^a	225 ^a	238 ^b	244 ^b
<u>Meat quality traits</u>								
Reflectance value, EEL	5.7	6.1*	1	8.1	18.6 ^a	18.4 ^a	21.8 ^b	19.7 ^b

*P<0.05; **P<0.01; ***P<0.001

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Table 1, continued

¹Only the traits for which the QTL peak was in the *IGF2*

5 region (0-10 cM) and the test statistic reached the nominal significance threshold of $F=3.9$ are included.

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²"QTL" is the test statistic for the presence of a QTL under a genetic model with additive, dominance, and imprinting effects (3 d.f.) while "Imprinting" is the test statistic for the presence of an imprinting effect (1 d.f.), both obtained at the position of the QTL peak. Genome-wide significance thresholds, estimated by permutation, were used for the QTL test while nominal significance thresholds were used for the Imprinting test.

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15 ³In cM from the distal end of 2p; *IGF2* is located at 0.3 cM.

⁴The reduction in the residual variance of the F_2 population effected by inclusion of an imprinted QTL at the given position.

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⁵Means and standard errors estimated at the *IGF2* locus by classifying the genotypes according to the population and parent of origin of each allele. *W* and *L* represent alleles derived from the Wild Boar and Large White founders,

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respectively; superscript *P* and *M* represent a paternal and maternal origin, respectively. Figures with different letters (superscript a or b) are significantly different at least at the 5% level, most of them are different at the 1% or 0.1% level.

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Table 2 Maximum likelihood phenotypic means for the different F2 genotypes estimated under (i) a model of a mendelian QTL, and (ii) a model assuming an imprinted QTL.

Traits	Mendelian QTL				Imprinted QTL		
	$\mu_{LW/LW}$	$\mu_{LW/P}$	$\mu_{P/P}$	R	$\mu_{PAT/LW}$	$\mu_{PAT/P}$	R
BFT (cm)	2.98	2.84	2.64	0.27	2.94	2.70	0.27
% ham	21.10	21.56	22.15	0.83	21.23	21.95	0.83
% loin	24.96	25.53	26.46	0.91	25.12	26.14	0.93
% lean cuts	65.02	65.96	67.60	1.65	65.23	67.05	1.67
% backfat	6.56	6.02	5.33	0.85	6.43	5.56	0.85
% fat cuts	28.92	27.68	26.66	1.46	28.54	26.99	1.49

Claims

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CLAIMS

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1. A method for selecting a domestic animal for having desired genotypic properties comprising testing said animal for the presence of a parentally imprinted quantitative trait locus (QTL).

5 2. A method according to claim 1 further comprising testing a nucleic acid sample from said animal for the presence of a parentally imprinted quantitative trait locus (QTL).

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3. A method according to claim 1 or 2 wherein in the pig said QTL is located at chromosome 2.

10 4. A method according to claim 2 or 3 wherein said QTL is mapping at around position 2p1.7.

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5. A method according to claim 1 to 4 wherein said QTL is related to the potential muscle mass and/or fat deposition of said animal.

15 6. A method according to claim 5 wherein said QTL comprises at least a part of an insulin-like growth factor-2 (IGF2) gene.

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7. A method according to anyone of claims 1 to 6 wherein in the pig said QTL comprises a marker characterised as nt241(G-A) or as Swc9, as identified in figure 4.

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8. A method according to anyone of claims 1-7 wherein a paternal allele of said QTL is predominantly expressed in said animal.

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9. A method according to anyone of claims 1-7 wherein a maternal allele of said QTL is predominantly expressed in said animal.

10. An isolated and/or recombinant nucleic acid comprising a parentally imprinted quantitative trait locus (QTL) or functional fragment derived thereof.

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30 11. An isolated and/or recombinant nucleic acid comprising a synthetic parentally imprinted quantitative trait locus (QTL)

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derived from at least one chromosome or functional fragment derived thereof.

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12. A nucleic acid according to claim 10 or 11 at least partly derived from a *Sus scrofa* chromosome.

5 13. A nucleic acid according to claim 12 wherein said nucleic acid is at least partly derived from a *Sus scrofa* chromosome 2, preferably from a region mapping at around position 2p1.7.

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14. A nucleic acid according to any one of claims 10 to 13 wherein said QTL is related to the potential muscle mass and/or fat deposition of said animal.

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15. A nucleic acid according to any one of claims 10 to 14 wherein said QTL comprises at least a part of a insulin-like growth factor-2 (IGF2) gene.

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16. A nucleic acid according to anyone of claims 10 to 15 wherein a paternal allele of said QTL is capable of being predominantly expressed.

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17. A nucleic acid according to anyone of claims 10 to 16 wherein a maternal allele of said QTL is capable of being predominantly expressed.

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18. Use of a nucleic acid or fragment derived thereof according to claim 10 in a method according to anyone of claims 1-9.

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19. Use according to claim 18 to select a breeding animal or animal destined for slaughter for having desired genotypic or potential phenotypic properties.

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20. Use according to claim 19 wherein said properties are related to muscle mass and/or fat deposition.

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21. An animal such as pig selected by a use according to claim 18 to 20.

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22. A animal according to claim 21 characterised in being homozygous for an allele at a paternally imprinted QTL, preferably located at a *Sus scrofa* chromosome 2 mapping at around position 2p1.7.

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23. An animal according to claim 21 or 22 wherein said QTL is related to the potential muscle mass and/or fat deposition of

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said pig and/or wherein said QTL comprises at least a part of
a insulin-like growth factor-2 (IGF2) allele.

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24. A transgenic animal comprising a nucleic acid according
to anyone of claims 11 to 16.

5 25. An animal according to anyone of claims 21-24 which is a
male.

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26. Sperm or an embryo derived from an animal according to
anyone of claims 21-25.

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27. Use of a sperm or an embryo according to claim 26 in
breeding animals destined for slaughter.

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FIGURE 1

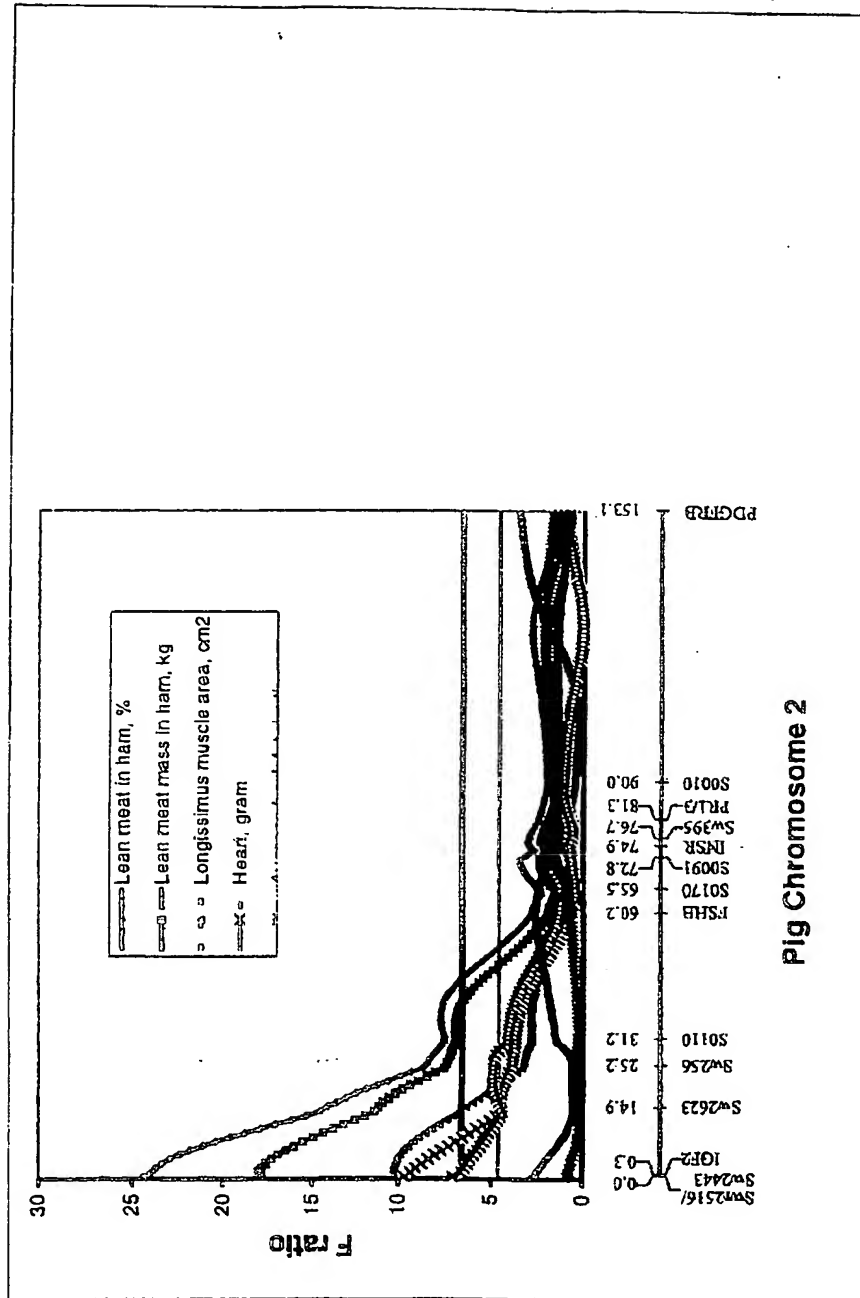
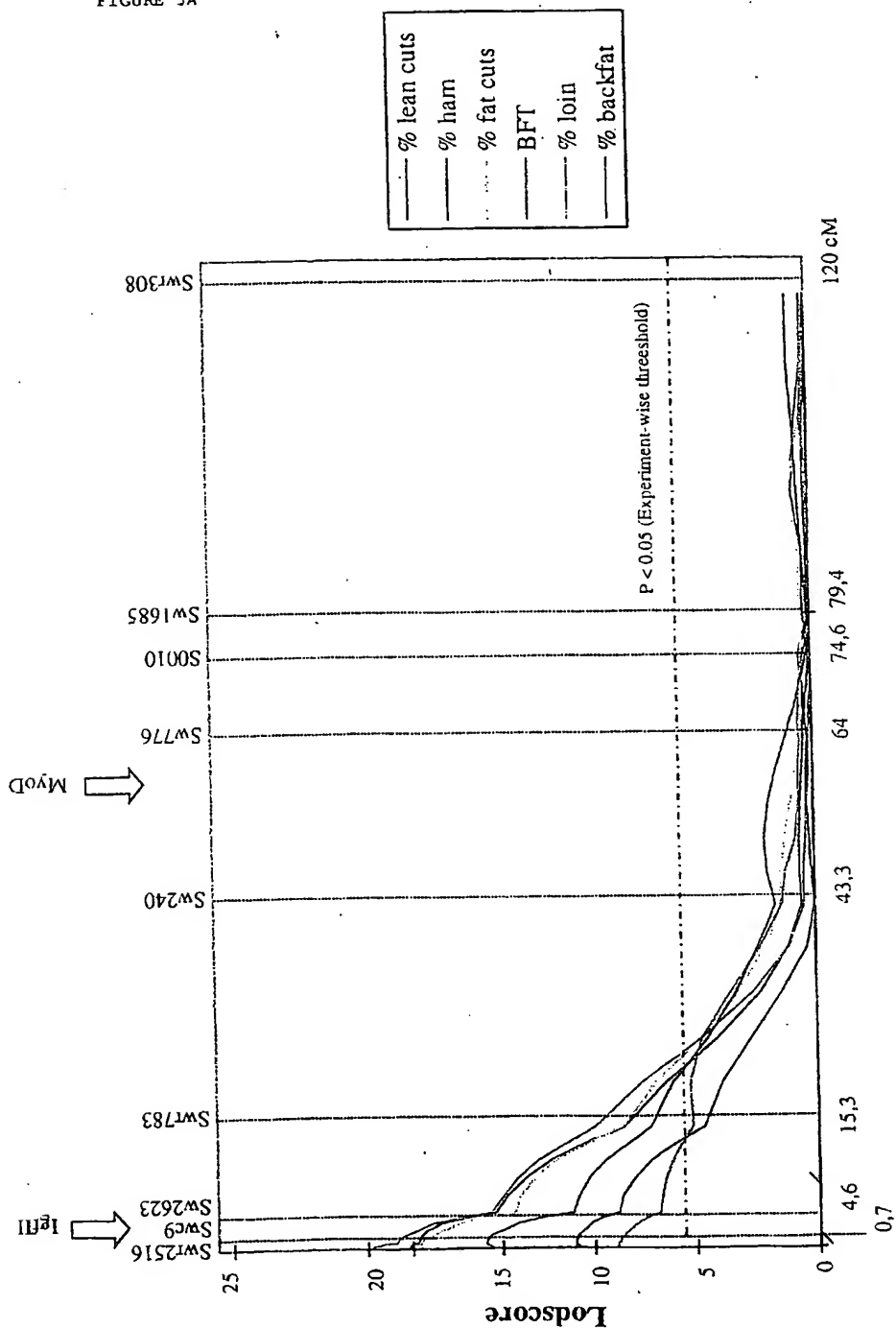


FIGURE 2

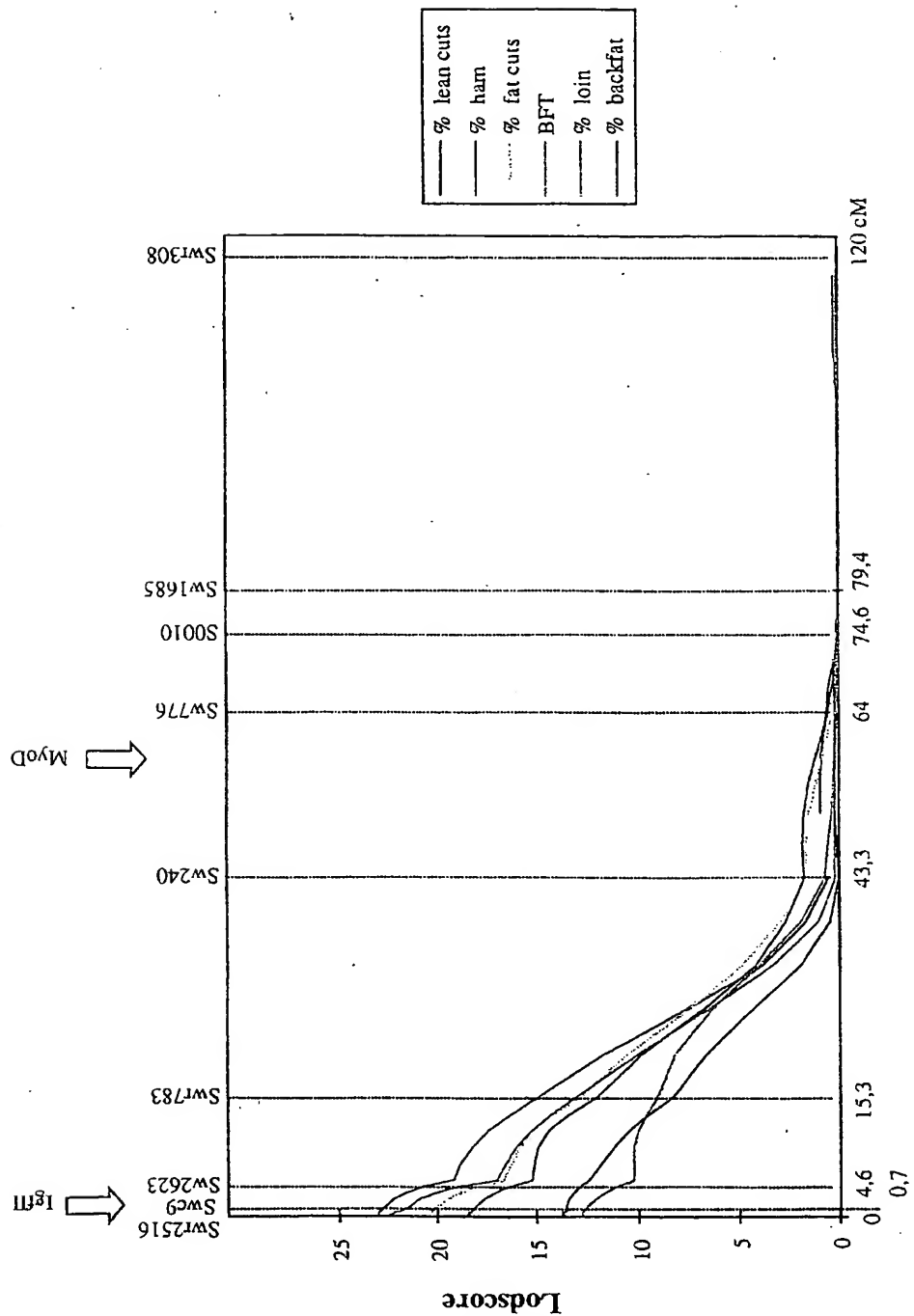


FIGURE 3A



(9) SUBSTITUTE SHEET (RULE 26)

FIGURE 3B



(26) SUBSTITUTE SHEET (RULE 26)

FIGURE 3C

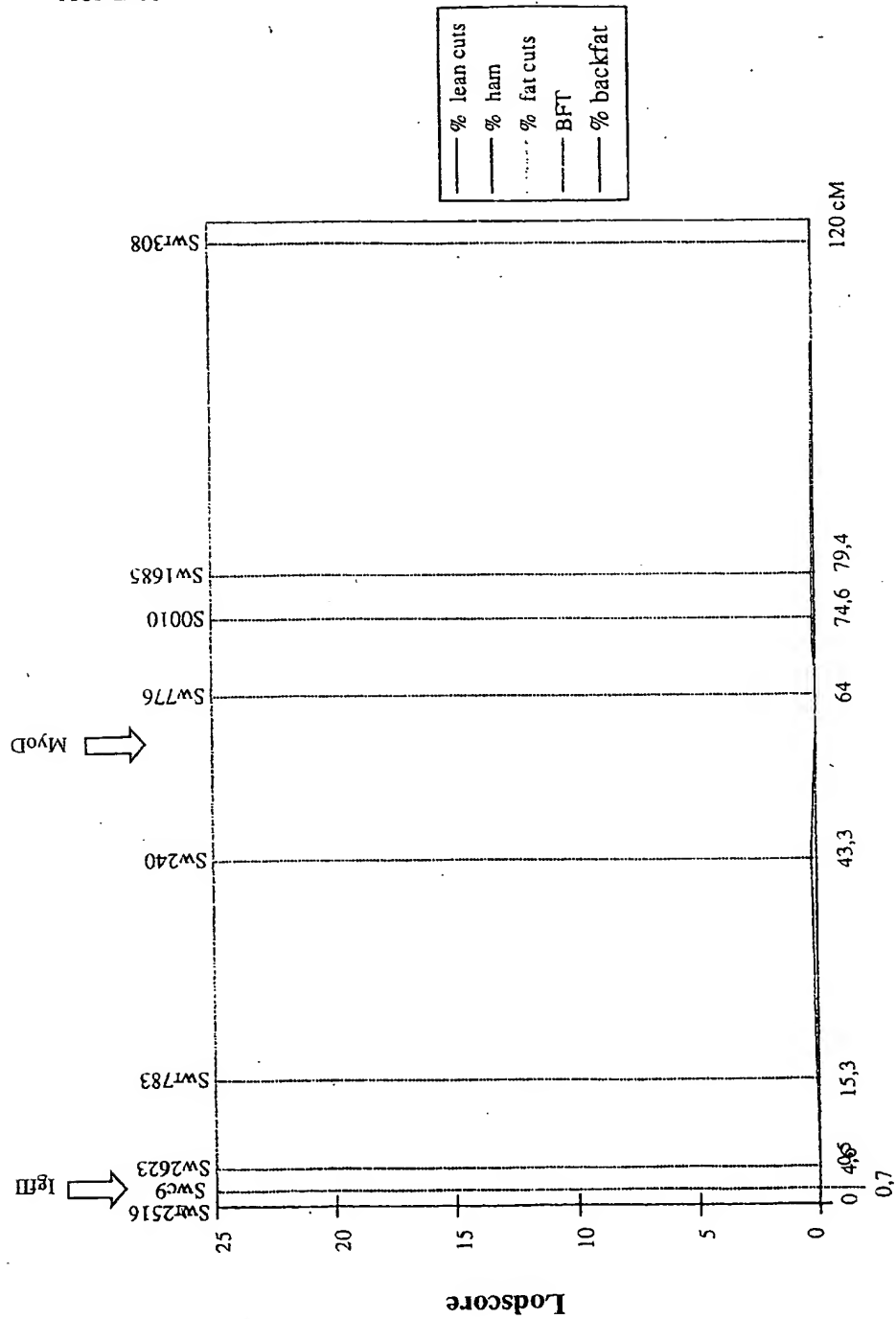


FIGURE 4

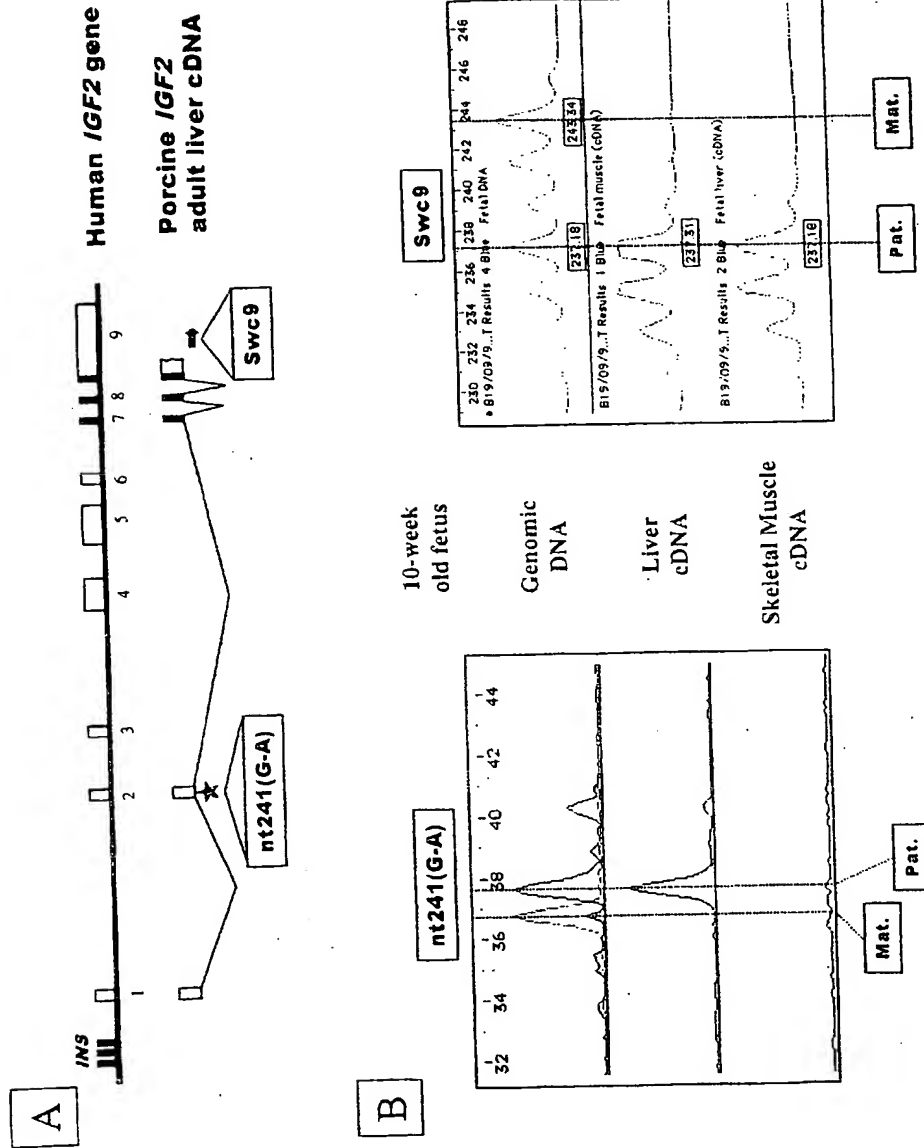


FIGURE 5

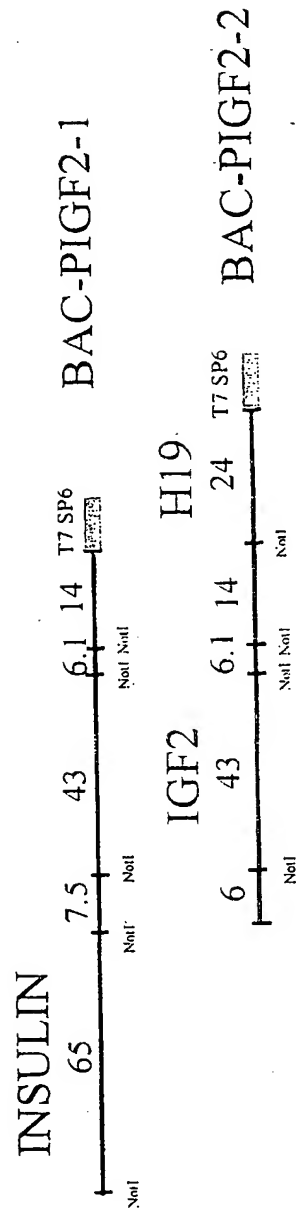


FIGURE 6

Contig 1 (500 bp)
GGGTGGGAGCTTCCCTCCAGACCCGAGGAGGCCAAGTTCCCTGGCCCTGCCACCCAGGGCCAGCTGAAGC
AGGTCAGAGACACCCGCTCCTGTCCCTCCTGTACCTAACCCACAGGCCGGGGCCAGGGACACAGGCCACA
TGGCATCTCCCCCATGCCCTGCCCAAGGCGCCAGCAGGTGAGGCTGGAGCAGAGTCTGGGTCTGCGGG
CCAGACCGAGGGCAGGACAGCTGGGCATCTGTCTCACAGTCCCGCGCTTTGTCCGGGAGGCGGACAGCCCTC
ATCCAAGACGCCCGCAAGGAACGGGAGAGGCGGAGGCCGGCTGCCGCGTCCGAGCCCGGGGAGGCCCTGG
AAGTGGGGGCCCTTGGCGAGCGGGACGGGAAGGCCCTGCTGAACCTGCTCTTACCCCTGAGGGCCACCAAGCC
CCCTCGCTGTTCGGTCCCTGAAAAAATTCTAGGTGAGGGGGCGGSCCAGGGCTCCCGGG

Contig 2 (943 bp)
TGCTCCTCACACCCCGGGCGGGGCTGCTCTTGGGGCCATCTCCCATGGGCCAGCACCCACTCTGGCCTTC
ACACCTGCCGTCTTCTGGGAAGTCCCTCTGGTTCCCAAGGAAAGTTTCTGAGCTGGACAAGTGCCACCACTGG
TCACCAAGTTCGATCCTGAGCTGGACCTGGACCACCCGGTGAAGCGGTGCCTCCCTCCCGGGCGCCATGTC
TCCCATCCCGAGGGTGTCCCACTCAGGGCCGGGACTGGGCGTGAACCCCGGGTGGGACGGAATGTTGGC
CTGCTGTGTGGCTCTGGCGGAACAGAGAGGCTGGCTGGGTGCCACCCCGAGGGCCCCCGCCGATGACACGG
GCCGCTCTGGGCTGGCGGGCAGGGCGGCCAGGC
AGGGCAGCCTCCGATGGGCTCCCGGCTGTCAACAGGCTTCTCGGACCAGTTGTACCGCCAGCGCAGGAAGC
TGATTGCCAGATCGCTTCCAGTACAGGCAGTAAGTCCCTCCAGGGCTCAGCCTGGGGGCCAGACCTCAG
CCTGGGCTCAGCCAGACCTGGGGTGGAGGGAAGGAGGTTCTCTTGTACCAAGCCACCACTTCACT
GTACCATGGTCACCGACTCTGGGTCCCAAAATCACAGCTGAGGAACTGGGSCACAGAGTGGTTAAGCATCT
TGCTGAAGCCACACAGCTGGCGGAATTTGGCCCGGCCCTCTCTGGGCTCCACACCTGCTCCCTGAGGG
GCCCGGACTGACAGCTGTCCCTCTCAGAGGTG
ACCTATTCCCGCGTGGAGTACACAGCCGAGGAGATTGCCACCTGGTGAAGCCCTGTGACAGCGGCTGGGAG
GGCGGGAGTGGGGGAAGGACAGGAAGACCTCAGAATTCCCGCTGCAACGTGGTGGCTCTATCATGA

Contig 3 (1500 bp)
GGGGAGGGGATGCTCAGACCCGCTCTGGGAAGAAGAGGCTCAGAGAAATCCCTTCCCAAGGTCACGCGG
TGGAGCCAGGGGCCCGCTAGCGGCCGATTCCCAAGCTCGTCTCCACCTGCTGGCGCTCCAGGAAGTGC
GGAGCGGTGGGGCCCTGGATGGGTCCGCGAGTGGGCTCGCAGGAGACCCCTGGAGGGGCTGCGGACACCC
ACCTGCCACTCACAAGGTGCCAAGCGCGGTTGGAATGAGCTGAGCCTCTCCCGCTCTCCCTCGGAGGA
CAATTGGCCTCGCATCCTTGGGGTCTCGGACGAGGAAATTGAGAAGCTGTCCACGGTGGGTTTCTCCCGTGC
AGGGCCTGGGTTCCAGCCAGGCCCTCTGTCCAA
GGGGTGTGCTCTCAGCTGTGACCCCGGAGGCTGGATCGGTTCTGCTGGGTGGGCGGTGCCGGGCCA
CGGGCAGCAGGGGCAGCGGTGGCGGCCCGAGCGGTGTCTGAGCCCTTGGCGCTGTCCCAACAGCTGTAC
TGCTTACCGGTGGAGTTTGGGCTCTGCAAAACAGAACGGCGAGGTGAAGGCTACCGGGCTGGGCTGCTCTCT
CCTACGGGGAGCTCTGCTGAGGCTTCCCAAGCGCTGGGCTGGGCTGGGCTGGGCTGGGCTGGGCTGGGCT
TSCCTTGTGGAATCCAGCTCTCGGGAGGCTGGAGCGAGGGGCTGCCCTCTG6GGGCAACCAAGAAAGTGGT
TSCGCCCCCTCTCACACACCTGTGCTGGGCGCTG
GGGGGACCCCTCTGGGGATGTGGTGCACAGCCAGGCTACCAAGGAGTCAAGACAGGGGCTCCCTTCCC
TCGGGTCTTGAACCCCTGGCTTCCCGCAGCACTCCCTGTCCGAGGAGCCGAGATCCGGGCTTTCGACCC
CGACCGGGCGGCGGTGCAGCCCTACCAAGGACAGACCTACCAAGCCCTCTACTTCTGTCTGAGAGTTTCAGT
GACGCCAAGGACAAGCTCAGGTGGGCGGGGCCCCGGCCCCCAACTGGAGGATCCAGCTGCAGCCCCGCC
TATGAGCCCATTTCCAGCAGAGGAGCTGCTGCGGACCCACCGCTACAACCCCCCTCCACAGCTGGAACC
CCAGAAAGCTTGGGAGGGGGGACCTGCAGGGCTG
TGGCAGGTGAGGTCAGTCCAGCCAGGCTTTTGGGGTGAAGTCTGACTTTGTAAGAGGGGGTGCAGGGT
CCTTCCAGCCTCTCCCTCCGAGCAGCTGGGGCGGGGCGGGGTGCGATGAAGGAGAGATGAGCCAGCC
ACCCGTTCACTCTCAGGAGGCGCTCTGTCCAGCCAGGCTCTGTGTGTCACAGGGAAGTGAAGGCCAGG
TGTGTGTGGGGGGTGAATCTCACACACAAGCTTAGGGACAGGACATAACGGCTCTCCAGGGCACACAG
TCTGGAGG

Contig 4 (3024 bp)
TTAANTCCANGTTGGCCCGACAAGTTTCCCCATTGAAAAGGGGCCAGTTAAGCCCCAACNCAATTAATTGG
AAGTTAGCTCCCTCATTTAGGCTCCCCAGNCTTTACNCTTTATGTTCCGGTTCGTATTTTGTGGGAATTGTA
GGGGATACAATTTCTCTCAAGNAACCACTATGCCCATGATTACGCGGTACAGTAGTTCACTAGTCCCCCGG
CCCATGGGACAGCGAAGGGAACCAATATGTCGTGGGGCGGGTCTAAGGGGTCAACACAGGGAGGGGAGG
GGCTCCAGGAGGACGGCCACTGAGCGGTACCTGGTGGGGGAGGTTGGTGGGGCCACACCCAGGAGTCTGTG
CCCCCCCCACTCCCGCGTTGGACATGAGAAGCAGGGGCCAGCTGCGGGTCCCTGAGTTACAGCCCCCCCC
CCCCACGCCGACAGCCCCGGGTCTCAGAGGCTGCTGTGCTGGGGGCGGGGCGCTTATGRRGCCGGGAG
CAGCCCCCCCCCAGGCTTACAGAGCATCTTGGGGCTCAGGGATGGACCGGGGCTGCRGGCAGGTGTCTCT
TCGGCCCCCACTCCCTGGGCTATAACGTGGAAGATGGGGCCCAAGCCCGGKCGTTTGGCTTTGTCCCCAG
CCAGTGGGACAGCTGGCCCTCAGGCGCTGTTAAGACTCTAATGACCTCAAGGCCCCAGAGGCGCTGAT
GACCCACGGAGATGATCCCGAGGCTGGCAGCAGGGAATGATCCAGAAAGTGCACCTCAGCCCCAGGCCA

FIGURE 6, CONTD.

TCTGCCACCCACCTGGAGGCCCTCAGGGGCCGGGCGCCGGGGGGCAGGCGGTATAAAGCCGGCCGGGCCAGC
CGCCCCCAGCCCTCTGGGACCAGCTGTGTTCAGGCCACCGCAAGCAGGTCTGTCCCTTGGGCTCCCGTC
AGCTGGGTCTGGGCTGTCTCTGTGGGGCCAGGGCATCTCGGAGGAGGACGTGGGCTCTCTCTCGGAGCCCT
TGGGGGTGAGGCTGTGGGGGCTGCAGGTGCCCTGGCTGGCTCAACGCCGCCCGTCCCCCAGGTCTCTCAC
CCCCCGCATGGCCCTGTGGAGCGCCTCTGCCCTGTGGCCCTGTGGCCTCTGGGCGCCCGCCCGGC
CCAGGCTCTGTGAACAGCACCTGTGGGCTCCACCTGGTGGAGGCGCTGTACCTGGTGTGGGGGAGCGC
GGCTTCTTACACGCCCAAGGCCCTCGGGAGCGGAGAACCTCAGGGTGAGCCGAGGGGYGTCCCGGA
GCGGTGGGGGAGTTTAAAGGAGAAATGTAAAGTGACCACTCCCTGGGAGCTGAGCCAGAGACACC
CTCCACAGCCCTGGTCCGCTCGAGAAGCCCTCTCTCCCTCTCTCCG
AGGCGGCTCCAGGAGGAATCTTACGGAGTCAAGGCCGGGTGCCGCTGGTCTCGAGTGACATGCCGCTGGT
GTCCCTCTGCCGCCCATATGCCCTGAGAGAWGCCCATCCCTTGGCAGGCGGCCCTGCCGGCAGGC
GGCGGAGGCGCCAGGACCGGTGGCTGTCTCGGCTTCACTCCAGGCTGGGCGGGGTGGGGGTGGCTGTCTCT
GTGTGACCGGCTCTCCCGCAGCAGGTGCCCTGGAGCTGGGCGGAGGCCCTGGGCGGCTGCAGCCCTGGCG
TGGAGGGGCCCGCCAGAGCGTGGCATCTGGAGCAGTCTGCACCACTCTGTTCCTCTACAGCTGGA
GAAGTACTGCAACTAGGCCGCCCTGAGGGCGCTGTCTCTCCCGCACCCCAAAACCAATAAAGTCTGAA
TGAGCCCGGCCGAGTCTGTGTCTGTGTGGCTGGGCGGGGGGCTTGGTGGGGGAGGGGCCAGAGGCTGT
GGGGGCTCTGCTGCGACCTCTCTGTCTCTGTCTCTGTCTCTGTCTCTGTCTCTGTCTCTGTCTCTGTCTCT
GCCCCAGGCACATGGCCACCGGGGACAGGGCCAGGGCAGGGCCCTTCAATGTGGCGAGCTCTGTCTCT
AGGGCTCCAGACACCCCTCTGGGTGCCACTGCTGCACAGGGTCACTCTGAGGCTCACAGGCGACCCACCC
AGACTGCTCTGGGCACACAAATAGCCAGGGCTCTTGGCTGTGTGTGTGTGTGTGTGTGTGTGTGTGTGT
CCCCCGGGGACCAAGACTTGGCCAGGCTGCCAGTCCCGCAGGCCAAACCAATCTGCACCTTTGCTGAAGGTT
CAGCCCGGCCAGCACTGGGGGCGGGCGGCTTAGAGCTGGGCGCCCGGGCGGCGGAGGACTGCACACCCGCGC
AGGTGGGCTTGGGGGTGGCAGCAGGCTCTCCGCTGGGACCCAGCCAGCTGGGAGCTCACTCTCAACACG
AGGCTCTCACTGTGTCT
GGCCCGCCCTTGGAGAGGCGCAGGGCTGGGCAAGGGGTGGGCGGCGGCGGCGGCGGCGGCGGCGGCGGCG
TAGGGGCTGGGATGAGTGGCGAGCTGTCCATGGGAGCACCCAGCGGCCCATTTGGCACCAGTACAGGCAAGG
GCAGCTGCAGCAGCTGAGTACGTGGGCTCCCGGACTGGTGGTGTCCGCTGCCCTCTGGGAGCAGCGGG
CTGAGCTTGTGGTCTTCAACCCAGGGAGACCCGTGACCACTCTGTGTCTCTCTCTCTCTCTCTCTCTCTCT
GAGCTCTTGGGACTCGGGGCGCCCTGAGCGGCCCACTCGCAGGACTTCAAGCGGTGTGGGCTCTGGGTGAG
TGGGGCTTGGGAGAGGCTCACTCTTGTCT
GGCTTGGGCTGGGGCTTCT

Contig 5 (1730 bp)

CGTCACCCGAGAGCCAGGCCACAGGCTTGGCTCAGCCCTCCACCCAGGCCACGTTCCGCCCTTCTG
GCAACTGGAGGACAGCCCGCCCTCGCCCTCGGACCTGGCTTCTGTTTGGCTTGGCATCTGGCAGTGGCGGCAG
CTGCGTTCAGCCCTGGATACACCTTGGCTTGGCGGTGGGCTCCCGTGTCTGAGGGCAGCCCCACACAGCT
CTGCTCACTTGGCTTGTGTCTGTCTCGCATCCCGTATCACATGCCATCTTGGGGACCCGTAGCGCTTGC
CTGTGTGGCACTGTGGCACTGTGTCT
TCTAAGGACATTTGGCGGTGACGCTCCCTCCAGG
CTGGCCCCCGGATTGCATCTGCTTCTGGCAGGATGAATGGCACCTCTGCTGACCAATTAGGGCTGTATTT
GCCTTCTCTGTTGGCAGTAAATATTTACTGTCCCTCCCTGTCTCTCCAGGCCGAGNCCAGTTCTTGGGGG
ATGGGAGGTGGACAAAGGTGCCAAGCAGCCCTGCTCTTGGGGCCAGTGTCTGTGGGGGCGAGCCT
GGGAAGGAGGAGCGAGACTAGGAACAGAGGCTGTGTCTCTGGAAGAGGCCCTGGCAGAGTTCCGGCTGG
TGTGTCTCAGCTAGGCTGTGAGTCTTCAACTTGGGAGCGCGGCCCTTGACCCAGCAGGCTGACCCCT
GGTGCACTGCTTCACTGGTGGGCACTGTCCCC

ACCAGGCAAGGTGGTCCGAGCGGTCAATCACAGACAGAACCAGCAGAGGGCGCCAAAGCCCCACTTTTGACAA
ACTCCCTTTCGCTTGGCCGAAAGTCCAGCGGCGAGGTGCACCTCTCTGAGGGCTCTGCCACCCCTGTGTC
CGCTTGGCAGCACTCACAGGGCTGGGGGGGTGCCAAGAGCGCGCTACCTGAGCTTGGAGGCGATGGA
GTTTAGGAGGAAAGAGGGGACTCTGGGGGTGACTTTCTTACGGCCCAATTTGGGGCCAGCAAAACGAGG
CTGGAGGAGGCGGGCAGCTGTGCCAGCTGGAGCCTTTCTGAGGGTCTCAAGGCTGGGGAAATGAGGC
TGGGGGCTGGGGGTGTCACTGTGGGCCAGGAGG
CCCCCTGCTCTGATTGGAGCCGCTCGGCCACTTGAGCCAGGAGGCTCACATGAGGCGGGGGCTGCAGGAGCA
GGACCTCGGGGCGGGAGGCTTGGAGGGGTCCAGCTGGGCCAGGGTCTGTTCTTCCCGGGTCCATGTC
CAGCGCCCTCCCGCTGTGGGAGGAGAGGAGGTCCAGGGCAGAAAGATGCGTGGGATGGGGGGGTGGTCA
GGGTCTGGGAGCTGTGGAACAACAACACAGCGAGGTCCTGGGGCGCCCGGCCCGCCCTCTCGGCA
CTGTTGTTCTGGCGGGGTGAGGAGCAGCGAGGAGATCTCTCGAAAGTGGAGACTGGCGGGGGGCCCT
CGGGTCTCAGCTCACCCCTGAGCTAGCCCGCC

ACTCGGCTCAACCTCCCGCAGGCCCTGGCAGGCTCTCAGGAGTCACTGAGGGTCCCCAAAGTGGCAC
CAGGAGCTGGGCTGGTCTGTACACCCCCACCCACCTCCAGTCTGAGATATG

Contig 6 (4833 bp)

ATGTGAGCTGCACAGCATGAGCCCTCGGCCCACTGCTGTGGCTTGGGACATTGAGGTGTGTGCGGCCAG
GGGACACACCTTGGCTCTCAGGCTGCCGTACAGAGCGGCTGGGTCTGANGAGGTGGGGGCTCTGGG
ACCGCTGGTGTGAGTTCAGGAGGGGGTCTAGCCACCTCTCTCTGAGAGTTGGTGGGTGGGGGCTCTCTTAT
CGTGATGACAACTGATTTCTGGAAGAGCCAGGTGTTTCTGAGGCTGTGGTTCACCTCTCCAGTGGCCA
CAAGGTGCCGGCTCGGTCAGATTGAGAAGCCCTGGGGAGCGGGTGTGATGCCAGATTACGCTGCTCT

FIGURE 6, CONTD.

[illegible]

FIGURE 6, CONTD.

CTGGTTTCGCACTCCTCCGGGGACTGTTGAAGTACCCGAGAGCGCNCGGGAGCGCCGGGGGAGCGGGGGTG
GCCGCCGGGGTGTCTCCGGGGCCCCGGACCGAGCCAGGGAGCGCTGCCCGCGCGCGAGCCGGGCGCGG
CTTCGCCTAGGCTCACAGCGCGGGAGCGCGTGGGGCGCGCGCTGCCGGAGTCCGCTCCTCTCGGAGG
CGGGCGACCGGGAGCCTGGGGACCCGAGCGCCGGGGAGCAGCGCCCGACACGCCCGGGCGCTCTCG
GCTTCCTCCTTCCAGCGCGCGCGCGCGCGCGCTTCCGACCGGGGCGCTCTCAGTGGCAGGAGAAGCG
TGGCTCCCGCGGGGTGGGGACCCGAGGAAAC
CGCACCGCTGGAGCGCGCGCGCGCGCGCGCTCGCGTCCCCGGGGAGGGCGCCACTGCTCCGCGCGG
CGTCCCCGACCGCCCGCGCGCTTCCCGCGCGCGCGCGGATCTTAACCTCTCTCTCGGTCCGAGCCCGCAT
CCCCAGGGCTCCAGGCCCGCGGACTTCCCGCTCCTCCCAATTGCAGACAGCACTTTTCTGGGACCTCCC
AAGGACAGCCTGGCTCCAGGGTCCCCAGATACATTACCACTTCTCCAGATCACAAAGTGGGTTTTCTGGGC
ACTAACTTCCAGAGCCTCAAAGCACATGAGCCCTACTGGCTTCCAGGTTTCCACTAGTGGCTCGGTCC
CCACTCACTGGGATTTCTTCCAGGCTCTTCG
GGTGTATCCCACTTTCGCGCGCGAGTCCCGCATGCCAATCCTCTCTAGAAAACCTTAACACTGACTC
CTGGTCTCGGGGTGAGGCTGCCAATGTGCTGACTCCCGAGAAGTATACCACTGTTTTCTGGCATTTGGG
CACCCTTCCCCAAAACAGTGAAGCTCTTTTCCCGCTCCCCATAATTTGGACCGCCAGGGCACCAAGCT
TAGCGCCCTGTTTGGCTCCCCACACCGCGAAGCCCTGCTCCCTGGGGTTACGACAGTTTGGGACTTTATC
TGCCAAGTTCACAACTGATTGGCCCCAAGCTGGGGTCCCTAAATTGTACACAAAGAACCCAGCCCCCCCC
CCCACTCCAGTACAGGAAGCGATGGCCCCAGGGA
CCCTCGGAGTTGGAAAGCTGGCTTCTTAAGCTTCCACAAAATTGAGGCTTTCGCGCATGGCGCGCTGATGCC
CTTGCTGAATCAGAAGCACTCTGCGCTCTGATTCTCTCTTCCACAACTGAGAGCATGATTTCTGTGCTCCC
CAACTCACTGAGCAAAATCTTTTGTGGGGCTGCAAGATAGGAGGCACTTCTCTCGGAGCTTCCAAA
CTCCCTTGCCTATATCAAGTTTCCCTAAAACCTTAGACAGAGCTTCCAGGCCCCAGAGGCACACAGGCCATT
ATTGGAGCTTCGTTTAAATGATGACAGGAGCAATGGGTTATGACAGTCCCCCAACTCACAAATGCCCGAGTAT
CCTTGGCTTACGCAAGCCCAAGCAAACTCTTGC
ACAGATCCCATATCTTGTATGTCAAGCGCTTTCGCTGTCCCAAGTAAACAAATAGTCTGAGTGTTTTCTCCAC
CTCATACATTCGGAATATTAATAAATTCCTGGGCCCCGGAGCTGACAGACAAGAAATCGGGCTTCTTAAA
ATTGCAACTGATTTCCCAAAATCCAGGCCAACGCCAGACCTCTCCCAATCTGGAGCCCTCCGACTGGACAC
ACTGGACTCTTAAGTATTACGCGCTGTCTCCAGGACCCCCAAATGCATTCAAAGTACGCTTTGGTCACAGA
AAGGCACTGATTTCTGGGCTCCAAAGCAGCCATGCACCCCGAGTCACCCCAAACTTAGTCAGCATTTCCG
GGGTTCTCTCCGCACTGCAAACTTCAAACTGCGG
ACACCGGTTCTTACGAGCCACCGCTAGACGGTCTTAATCCCTTTTCCCCAGACCTAGATTG
Contig 8' (371 bp)
AGATTCAAAACTATTTTTCTGGGGCTCCAAATTGAGGTGTGCTTCCAGTCTCTCAAAATAAACTGAGGG
GTTTTTGTCTTCTTTCTTTTCTTTTCTTTTCTTTTCTTTTCTTTTCTTTTCTTTTCTTTTCTTTTCTTTT
CCATTGATTTATGGTCCCTGACTTTATGACCTTGGCCCCAAGTCCCTTAAATGTAGGCCATTTTCCAGCG
GCTTCCCAAAATGAATTTGCCAGATCCCGCGCAAAAAATATCCCGGGTCTTGAAATGTCAGGATTATACA
GGCTTGGGCTGACACCCCTCTTCTACTAACCAGGTTCCCTGAAGTTTAGAGATCACTACCTAATGAACAA
ATCCAC
Contig 9 (2415 bp)
CCAAACTGGGGCCCTATCTTACTAGGTTCCCTAAATGCAGACAGCGCCCGGAAATAGGGGCGTTTTTTT
TCTGTGTTGCCAAAATAAATTAATTAAGCAATTTTATAGATTAATCTAAATGACCTTGATTTTCTGTC
GTCTCCAAATGTACTTTTACAGCCCGAGTTGCCCTCAGTTTAGACGGTGTGCTTGAATCTTAAGCACC
CTGAGGATTTTCCGAGGAAGCCACCAACTACGGAATTTACTGTCTTCCGGGGCCACAAGCTCCAGGCT
ACCAACTTGGATTTCTAAACCGTGGAAATCAGGCTCCACTTCCCTCCGCCACCCGAGGCTCTGCTCAGACCC
CCCAACGTTGCCGCTGTTCTTCTCCCCCAAAAT
TTATTTAGAGAATATGCTCTCTCGGTTCTGCCAAGTTTTCGCTCAGACTTCTCGGTCTATCCCAAAATCC
TCTTCCCAAGTCCGGGAGCCCCACAAGCTTACCGACCCACATGCTGGGTCCCCCAACTTAAACGCCATC
CCCTGTCCCCAGATTCACCGAGTGAATTTCTGCTGCTCAGACTGGGACTCTTTACTGGAGTCTCGAATTT
AGCCATTAAATCAGTTCTCCACTCCGACGCGAGGCTCCCTTGGGTCCCAAGTCCGGGACATGGGTTCTCTTG
CTTGCAAAATCAGGCTGCTGACTTGAATTCAGGCTTTGGGCTTGGGCTTGTCTCCCGCGCGCGCGGCTCTGCTC
TCCCCCATCCCGCGCACGACGGCACTGGGTCTG
GGCTCTTGTGTCTCTACAGTCCCGGAGCTCCTCGGACTTGGGAAGTGTCTTCTGCTTCCCCAAATAC
ACTCGGCGCGGAGTGTGTCGCCAGGAGTAGGACAGGCTTCTCCCGCTCCAGGAAACAGACTGGGCTTGG
GGAAAACGCTTCCGCGGCCCAAGCGGGGAGGACGGGCTTGGGAAGGAGGGACACGCGGAGAGGAGCAGC
CCGCTGGGCGCGGAGCGCGCGCTCCAGCGCGCGGCGGAGGATCCCGGAGGCGCGCGGAGCGCGG
CGGAAGTGAATGATGGCGGAGCGAGGGGCGAGCGGATCCCGGCTTCCCGCGCGCGCGGCTTCCCTCTG
GAGGACTCGGGCGCGCGGGTTCTGGGGCGGG

CGGGGCGCGGGGGCTTGTGCGTGCTCCACTTGGTAAAAATCACAACTACTTTTACGTGCGCCCGACTCTC
CAGGAGATGGTTTCCCCAGACCCCAAAATTCGTGGTGGCCCCGGGGCTGAACCCCGCTCTACGCAAGGCC
AACCGGCTGAGGACGGGGGACCAATTCAGTATTTTGGGTGGCGCCCCAAAGCGAGCTGCTTAGACGGCC
CCCGGTGAGCTCGGTCTCGAGCTAGGCTTGAGCGAGGTTCCCGCCCTCTCTCTCTCGGGCAGCG
CGCGCAGGCGCGCGCCCTCCCACTGACGGCACTGGCGGCGCGGAGACGACTCCCGGTTCCCGCGCGG
CACCGGGGGCGCTTGGGCTCTGGCTCGCGCTCGA
GGCGCTCGCCCTGCTCGGGCAGGTGAGGCTTACGCCCGGGCCCGCGCCAGGACGACCCCTTACCCCGCAG
CTCCAGCGGGACTCGGGGCCCCGGATCGACGCTTACGCCACTGTGCCCGCACCGCCCGAGGGGCTGTGA
CACTTACCACCTGGCGGCCCGGTCCCCCGCGACGAATGTAGGATCTCTGACACCCGGAAGCTTAAGC
GGGCGCCCCATACATTTCTGTACGGATTCGGGATTCTCTCGAACTCTCGAGATCTGTATGGCAAGGTGA
TGGCTGCAATTATTTCTGATAATTCAGCGAAGACTGCGCACCAGAGCTATCGCGCTCTGGGTTTTAAGGC
GAAACCCAAATTACGATCTGGTCAACGAACAT

[illegible]

FIGURE 6, CONTD.

CCACACACACATGCATTCAACACACACACACTCGTGCATACACACGTGCGCGGCACACACACACACA
CACACTCTCTCTCTGTGGGATCCCTGAG
Contig 19 (500 bp)
TGGCTCTGGCATAGGCTGGCAGCTGCAGCTCTGACTGGACCCCTTGCCCTG
GGAACCTCCATATGCCGTGGAGCGGCCCTAGAAAAGGCGAAAAA
AAAAAAAAAACAAACCAACAAACAAACAAAGCCAAACACACAGAACTC
ACAGACACAGAAGAGACTGGTGGTGGCAAGGTGGGGTCGAGGGTGGG
AAAAATGAGGAGAGGGGGCAAAACACAAACGTGCAGCCATAAAATGGT
AAAGTCCCGGGGACCTCCGGTAGCGCGTGTGGGACTCGGGTTGAGAACA
CACCGTGATGTGTATTCCCGAGTTGCTAAGAGTCCCTGTTGGAGAAACAA
ATGCGTATCGACGTGTGGAATGAAAGTTAACCCGACCTGCTGTCTGAT
CACTTTGCAACACATACAGACATAGAATCATTATGTTTACCCTGGAGC
TGACAGCGTATACGTCCCCAGCCTCAATTTAAACAGCGTTGCGGTG
Contig 20 (400 bp)
TTCATACTGTGCAATGCCAGCCCTTAAATGCACAGAGGAGCATTAACCTT
CTTTCCAGAACTCACTGAAATGATACCACTCATGTTTGGCACTTGCACTT
GGGCGTTATTTTATTGGTGGCGGAACAGCGCGATGTGGCACCAACTAG
CGCCGCTGTTTTATTTCCTCGGTATCCGCGCTCTCGCTGTCTTCCCC
CCCTTCCGCTTGAGCTGAGGAAAGGGCTGAGAGGAGGAAAGTCTGCATT
CACCCATCTCCCTCTGCTGTGTCTCTTACAGAAAGTGGTGGCCT
GTGCGGGGAAGTCACTAAACCTAGGACAGTGTCCCGTGGGTTCATGCTTG
TTACACCTTTGTGCACCTGGCCCAAGTCTCGGTGGAGCGAGAAGCTGGC
Contig 21 (559 bp)
AGCTAGCCCCCAGCCAGGGCCAGGCTCTCTCCACCCGCCCCAGCCA
GCATCTCTCAAGAGGAGGGGGCTCTAAGGGATGAGGACCTGCTCCAGTC
GGAGACACGAAGCCCCGCGGCTCTCCCCGAAAGTCCAGCTGCGGCTTT
CGAGCAGGCTGCGCCCTTCGTCAATCATTTCAGCCACAGAAGTGAAGG
CGCTTTCGTGGCGAGGCGAGCGGGACAGAAATGGAATCCACCCACAG
GCGAAGAGCCGCGTGGGTGAAGCGGCTCTGTTGGGACCGGGCGGG
AACTTCACATGGGGTCTGCTGTCTCCCATCTCCCATCGTCATTACTGCAG
GGGCTCGGCCACACCCGAGCTGCGGGGGCAGTGTGACACTGGACCT
GGCTTCGCTCCTATGATGTCTATGGGGCGGGGCCAGCAGGGCAGTGGC
CACACCTCGGGCTCCAGCACCAGCCAGGATGCGAGAGGGCCCCACCC
ACCACGGGCGATGTACATCCAGAGGACAGCTGAGCAAGGCTTGATANG
GGCTTCAAC
Contig 22 (450 bp)
CGTGCAAGGACCCGTGCGGGCTTCTGTGGCCACAGACAAACACAC
CATTATCTTACAGCCACCGCGCGGCTGTAAATGGGTAACCTGGGGCAA
GGGGGCCCCCTGCTGAGGCGGGGTGGGGAGCGCAAGGATGGCTGTGT
GCCCCAGCCAGTCTTTCAGGGCGCTGCTGTCTGCAACCGGCGCCCCAG
GAAGCAGAGCACCAGCTTCTCCCTATTCTAGAACCAGCCCCAGAAC
CTGGACCCAGACCCAGGCCAGGGGATACTGACAGAGCCAGGCAAGGG
GCCACTCCACACCCACAGAGGGGCCAGCAACCCAGTCACTGCGCAGC
CCATTGCCAGGGGGCAGATGGACACAGAGCAGCCCTCATCCACAGCAG
GCAGGGGAGTGAACCTGGTGCAAAACGGGGCGTTCCAGGAAGTTAAGCA
Contig 23 (535 bp)
TGCCAGAGACCTCAGAGCTGGGCTCTGCTTCCGGGGCTGACACGGAGG
CTGTGGCTTCCACACCTCAGGCCACAGCCAGCTGCCAAGTCCCTGAA
GTGTCCCCAGAGGTGSCCTGCTCCACGCCAACATCAGGCTGCTGCA
GCCCTGGAGCGGCCCCCTGTCCCGGSAAGCCCTCGGGCTCTCTCGGTC
GCCTCTGGGGAACCTCGGTAATGTGGCCAGCCGTGCAGTGGCCGGAT
ATTGCTCAGGGGGGCCAAGGCAGGGGGTGACACATCCGCAAGTACCG
CATATGCACAGGATATGGATTGGGTGTGGATTTAACCTTTTCGCAATGT
CTCTGCCGCTACAAATATTGTTTCTAATCTCTGCTCCCTGAGCCGCTG
AGTCTGCCCGGGAGCTGCGGGGAGCTGGCTGTCTGAACCTGCCCTGGCCC
CCACCCCAAGGGAGCCCCCGGCCAGTGTGAGGGCAGGAAGCTTGGGCA
CAGGCTGCAGAGGCCAGGCTGGCTCACTCACCT
Contig 24 (868 bp)
TATTGAAGACCTATCATGAGTTCCAGAGCGGAGGGGTGGAAGCAGGG
CCTACAGCCCACTCCCATCACTCCAGACCCGTCCGGGGCTGGTGTCCCC
TGCCCCCTACTCTCTCTGTTGGGGGACGCTCGAAGGAGGCACTCTE
GCCTGGAGCCTGGAGGGTCCCTGAACCTCCGCTGCCACCTGGGCCCTCG
GCTCTCTGCGCTGGGACCCGCGGTGGTGGGAAGCAGCCCTGCTCAGTG
GGAGGAGCGAGGCTGTGGCCGCCGCCACGGCCCTGGGGGGACGCACG

FIGURE 6, CONTD.

CAGGACGCANGTGGGCGTGTGTAGTCCGTCTACAGTCCAGCCAAGGGC
GGCCGCGACCGGCCAGGGTGGGCAGCCCCAGCCTCAGCAGGGCGCTCTCT
GGGGCTCAGGCTGCGCCGACGGGAGATGAGGGGTGAGGCGCAGTCTGGGG
CTGCTGCCCGCAGAACCTCGCCACAGTGGCAGCTGGGCACAGGGAGACCTG
TACTCCAGAACCTGAGGCTGGACGTCCSAGACCCGCGTGCCGGCCTCTT
GGTGCCCTGGTCAGGGTCTCTTTCTGGTTTGTGGGCAGAACCTCTCAG
CGCGTCCCTTGCATGGGGTGCTTAATCACGGAGTAAGGAGCCAGAGAAATGAG
GCACGGAGTATCCAGTGTAAACCTGGAGTATGGAGACGGGAGTACTAAT
TGTGGAGCATGGCTCTAAGGAATGGAGTATTCGTACGGAGAACCGGGG
CCGGGTGAAATACGGAGAGCGGGCTACGSAACAACGGGGACGGGGTATCCG
AAGGGGAGGATGGAGTATCGSCCGGAGGGTGGAGAATGGACACTAGAGGA
TGTATANNNGGCGTCAAT

Contig 25 (500 bp)

ACCAGTTTCGATGAGCAATCCCAGCGCGTAACATTATGGCTGCAGCCTG
GTCAATGCCGGTGGAGTTTGAACCTCCACGCGTGGCGATTCTGGTAGATA
AATCGACATGGACCAGGGAGTTGATTGAACATAACGGTAAATTTGGCAATC
GTTATCCCGGGCGTTGACGCACTAATGGACGTGGGCGGTGGGAAGTGT
GTCGGGCGCTGATGAAGATAAATTTAATTGCTATGGCATTCGGGTTGTGA
GAGGCCCGGTATTTGGTTTGCCTCTGGTGGAGGAAAAATGTCTGGCGTGG
ATGGAGTGTGATTGCTACCTCCGACTTCTCCGCAAGAAGAATACGACAC
GCTGTTTGGCGAAGTAGTATCAGCAGCGGCAGACGCACGGGTATTTGTCG
AAGGCGGCTGGCAGTTTGATGATGATAAGCTCAATACGTGCATCATTTA
GGTGTGGGACGTTTGTACCAAGCGCAAGCGTGTACGGCGGGTTAAGC

Contig 26 (900 bp)

ATGTTTGATGTCCGCGGTGTGTAAAAATTTACGCTGCTCGGCTTCTTT
GGCTTCGTCCACCACCGGAAACGGACAAAAATTTCCGTCATACCTTTT
CTTTAGGCGGAAGCCAATGTCGTAATCTTCAGTAAGACTCTGCACGTG
AAAGCAATACCGTCACCGTCAGCTAACAGTGGCGTCAAGGGCGCGGCT
GAAACAGGTGCGGACGCTGCGCTGGGCACTTGTCCGGCGAGGGCTTCAC
GCACCGGAACATCTTTGCCATGCGAGCTCTGAAAACCTCATCAATGTAAGTC
ATGCTGGTGAAGTGGCTCCATTCGCGTTCGAACGGATACACGGGATCTC
AATCAGATCTTTACGCTCGACCAAGATGTTGAACAGACGCAATTCATCG
GTGAAATCACATCTTCGGGTCATGCAATAAAACCAACCAAGCGGAAA
TTGGCGCTACGCTCAATTTGGGTGATGGCGTCCAGCAGCTTGTTCAGACA
GTCGGCTTTGCTGGTGGGGCCAGGACGCGCGCAGACTACCTATGCAAT
TCGGGAAGCGAGCGCACACTTCGTCAACATCACGCTGAGTATCGGGGTG
TTGGGGTAGGTGCCAACAAAGATATGATAGTTTTCGTAGTCGAGCGTGGT
CGCGCGCAGCTCGGCCATATTGCCGATGACGCCGTTTCATTCCACGCCG
GAACCATAATCGCTAACGCTTTTCATCTGGTTTATACAGTTCGGGGTAA
CTCATTCGGGGTAGCGGGGATAAACACTCAACTTGGCTTAATGCGGGC
TACCCAGTATACGACATCTATAAAAAAATCGTCCAGCCGCTGATGAACA
TGATACCGCTAACCTTATCGCGATTACTTTAAGCCGTATAGCCAGGTA

Contig 27 (500 bp)

AGCTGGATGCCCCAGCTGTGGTCCCTTCCCTCAGGGCAGGTTCT
GTCCCTCTTGCAGCCACCGTCACTGCTGTGGACAGGTCTGCACACCGGCC
GTCCACCAAGAGCGTGGCAGCTCCCTGGGCACCGGCGCGCTCTGACGCA
CCATGTGTTCAAGGCAAGAGCACTGGACAGAGGGTCCAGACGTCCCTTG
TCCTGCTCAGGCCTGGGCGGGGGCAGCCCTGGCGGGAGAGGCCCTGGGCA
TCAGAGCCTCTGTGGCTGGAGCTTGGCGCCCTGCCCTCCCACTCCGT
CCTGCTCCTCGCGCGCTGCACGGACCTCTCCCGGCCCGCCAGGCTCATT
ACTCTTAAGGACCCTAGCCCCCTATGCTGAAATGCTGTACCTCGTGCTTG
TTTTCATCTGTTTATTACCTTATCTTCAATCCTGCTTGATGATATCTGGT
TATTTCTTTATTGATTATATATATCTTGTCTGTGTTTTATAGGACACTGT

Contig 28 (450 bp)

AGTGGGTGGGCGCTCCTGACGCTCAACACCGTATTTCCACGCGACCGC
GGATTCAACCTGCTCACACGGACGCCATGTAGACATGTTGGGGTTACGC
GCAGAGAAGCGACCTGCTCAACCGGCTGGTGAGTCGGGCGGCTTCGCCC
AGACCGATGGAGTCGTGGGTGTAACCATCACCTGACGCTGTTTCATCAG
CGCAGCCATACGTACGGCGTTACGTGCGTATTCACGAACATCAGGAAGG
TGGAGGTGTACGGCAGGAAGCCACCGTGCAGGGAGATACCGTTAGCAATC
GCGGTATACCGAAGTCCGGAACACCGTAGTGGATGTAGTTACCCGACG
ATCTTCGTTGATTGCTTTAGAACAGACACAGGGTCAGGTTAGACGGCG
CCGGGTACGACAGAACCCGAGGAATTCGGAACAGCCGACGACGACGCT

Contig 29 (450 bp)

SUBSTITUTE SHEET (RULE 26)

FIGURE 6, CONTD.

TCAGGCCAATCTGTCTGGTCTCCAATGGGGACAATTGGGTTCTTTAGGCT
TCTGTCCAATGGTCCGAATGGCCCACTCCCCGGGCGCCGCCAAGGGTCC
TCTGTGCCTCGGGTGGGCTGGCACGGACCGCCCCAGGGTCGTGCCAGCC
CCGTACCGGGGCCAGAACTTCGGGCTCTAGCTGGCTAGTCCGGGCTG
CTGTGACGGGGGCTGCGCTGGGGGAGAGGCGGGGTGAGGTAAACCTC
CCAGCCGCCGGGTCCCTGCCGAGCCCTAGGCGCCGAGACGGTGGCTG
GGTCGGTACCGCCAGACCCGAGGGCTCGGGGCCGGGTGACUCCAGCTG
TCGCACACGCTCGCAGCTCTCTTGCTCATAGGGCTCATCCCTCTGAGC
TCTCTACTGCCCCACCTCACCCGCTGGACCCCATGAAGCCCCGCGGA

Contig 30 (600 bp)

TAAACTAGCTCTAGTAGAAACATTATTTAAAAATAAAAAACCTGACT
ACGTCGGGAGTTCCCGTTGTGGCTCAGTGGTTGACGAATCCGATGAGGAA
CCATGAGGTTGCGAGTTCGATCCCTGGCCTCGCTCCGTGGGTGAGGATC
CGGCGTTGCGGTGCGCTGTGGTGTAGGTTGCAGATGAGGCTCGGATCCTG
CGTGGCTGTGGCTCGGGTGTAGGCGGCGGCTACAGCTCTGATGAGACCC
CTAGCCTGGGAACCTCCACATGCCCTGGGAGTGGCCCTAGAAAAAGGGCA
AAAGCAAAAAAACAAGAAAAAGAAAAATAAAATAAAAAAGACTATGT
AAATGAATTAACGACTGCCTAGGGTGGGATTTACAGCATGGGAAGTACA
GCATGGCCGTGACAGTCCAGGGTGAAGCGGGAAAAAGGAATAGGTTAG
GTAGTTTCTCCTGCTATTGTGATGTGGTCTGCTATCGCTTGAAGACGG
ACTGCACTGAGATAAATATGTACAGTACGATCCGAAAAACCGCCAGAAC
GGCAAAACGAATGACTCCAGTAAGAACCCAAAAGAGAAAAAGGAATAAT

Contig 31 (450 bp)

GCGGGGCGTTCCGGTGGGGTATTTAACTGGTCACCGSTTCGGCGGGC
GCGGTGCGTAACGAATGACCACTAACCCGCTGCTGCCAAACTGTCTGT
TACCGSTTCGACCGAATTTGGCCCGCASTTAATGGAACAGTGGCGAAAG
ACATCAAGAAAGTGTGCTGGAGCTGGGCGGTAAACGCGCGTTTATCGTC
TTTGACGATGCGGACCTCGACAAAGCGGTGGAAGCGCGCTGGCTCGAA
ATTCCGCAACGCGCGGCAAACTCCCTCTGCGCCAACCGCTGTATGTGC
AGGACGGCGTGTATGACCGTTTTCGCGAAAAATGACAGCAGCAATGAGC
AAATGTCACATCGGCGACGGCTGGATAACGGCTCACCATCGGCGCGCT
GATCGATGAAAAATCGGTATCAAAAGTGAAGAGCATATTGCCGATCGCG

Contig 32 (450 bp)

GGTGGATGCTGGCGATAGCGTCATCCTCGCTTATGCCGTGCAGCGGGCAA
GGATAAAGCGCGGATAAACATGACCCGGCATCAGCCCCATGCCCGCAGA
GTACGGATTACCTTGCCGCTCAGCGCCAGCGTGTAAATGCGTGCGCCGT
GATACGGCGCGCTAAAGCGATGGTGCGCTACGTTTSGTGGCGCGCGG
GCGATTTTACCGCTTTTCCACCGCTTCGGAACCGGTCTAACAGCAG
CGTTTCTTGGCGAAATCGCCCGGCACCTTCTGATTATAATCTCGCACA
GCTCCAGATACGGCTCGTAAGCCAGCACCTGGAAGCAGGTGTGCGACAGT
TTTTCAACTGCGCTTCACCGCGGCCACCCTTTCGATGCAAGTGCCC
GGTATTGAGCACCGTAATCCCGCCGCGAAATCAAGATACTACCGGCTT

Contig 33 (500 bp)

ACGTGAGGTTTGGGGAGGAAAGCGGGGACGAGCAGCCGAGAGGAGTG
GGGGCTGCGCTGTGGCTGATGAACTCTGAGAAGGTTAAGAGCCCCATT
TTTGTCTTCTCTTTTATATGGAATTTCCAAATGGATGCAAAAGTC
CCAAACCTAACTGGACATCTCTTGTACAGGAACGGTCAGGCACTTAT
GATGACCCGAGCCCGAGGGAAAAACCTGCGGTCTGGAGCCACCGGTC
CAGCAGGGCACACAGGCCCGACCCGCAAGCGGCAGGCTGAGTCAGTGA
ATGGCGTGCCCTCTGGTCAAGGACGGGCACTCTGGACCCAGGGAAGCCT
CTGAGGAGCCCTTTCACAGCGTCAAAACTGTTAACAGSGCCATGTTCC
CACCCCCCACACAGTGGTTGAGAAGCAGACCCAGGCATCGTAATATG
TCATCCGTGAGTTCCCTGTGTGCCACCAACAGAAAGCCCATCGTCACGTT

Contig 34 (400 bp)

CGGCATCGATGTACATGGTACGCAAGGCACTCGTAAGGCCCGAGCCTCT
AGGCCCTTCTCATTTGTCAGTCTCTCGCGGGGATCAGCAGCCAGGCTTG
TGACCCGGCCACTTTGACAGATAAGGACACAGAGAGGCCACAGCACTGG
TGTGAGGCCCCACAGCCAGCAGCCAGGGCAGGGAGGACTGGGTCTCACC
TGCCCTCAGCTGGGCCCAGCCTCCCTGGGAGTCCCGGAGTCTCCCCAGCTT
AGGAGTGTCCCTGGAACCTCTTCTCTCCCTTCCGCGCTCACCCGAC
CCCTGCTCCCCCAGCAACCCCTCCCCCTCTTCTTACCTTGAG
CTCCCCCTCTGAGGACCTCTACTGTTCTTCTTATCTCCCTTTGAGCCA

Contig 35 (500 bp)

TGGCGGTGAATATGTCTGCGTGAAGAGCATTTGTGGTGGTAGCGCGT

SUBSTITUTE SHEET (RULE 26)

FIGURE 6, CONTD.

TATATGCGGGAAGTTTAGGCGAACTGGACAGCCTGGGTTTATCCGGTAGC
GAAATCCGCTTTCACGGTAAAACGCTGCTAGCGCTGGTGGAAAAAGCSGA
GACATTGCCGGAAGATGCCCTTACCGCAGCCGATGCTTAACCTGATGGACA
TGCCGGGTATTCGTAAAGCGTTTAAAGCGATTAACTCGCTGATTACTGAC
GTGAGCGAAACGCATAAGATCAGCGCCGAATTGCTGGCATCGCGTCGGCA
AATCAACCAACTGCTGAACTGGCACTGGAACTGAAACCGCAGAACAAAT
TGCCGGAGCTGATTTCCGAGCTGGCGTGGTGAGCTGATGGCGGAAGCATT
ACACAATTATTCAGGAATATCCGAGTAAATCTTCCGAAGCCGGAAT
GGGCGCGCTCAGCGCCACATCCGGCTTCGGCAACTACAAATCCAACACC
Contig 36 (500 bp)
GATTTACAAGCCTGACCCACGCGAAATGCGCTAACAGCGTAAAGTCGT
GCGGCCAGAATTTTTCTCTCTCTGCTTTGCGCTCAATTCAAAAGTCAGC
GCTACGCCATCAGCATCTTCATGATGTGATTTAGCGCTCCACGGCAGGTT
GCGGGCAAAACCGTGGCAGGCAGACCTTGTGTGCGCGCGGACCAACC
ACGGCCAGCAAAACCGGTACGCCACCGCGAATAGCGACGCCATTTTGAAC
GGTGTGTGTGCTCAACACAGAACTTCTTCTCACCGCAGGTTTCCA
CGAGAGAAGGTGTGCGCCCTGTAAATGCAAAAGAGGCTTTTACTGGGAT
GATCCACCACAATGAGGTCCAGTTCATCCAGTTTACGACGGGAGAGGACA
GGGGAGATTTGTTGATGACCGGAAGGGCAAAATTTCTTAATCATGAC
GCAGTCTTTAACTTCATTTTATCAGSTAAAAAAGAGCGACCGAAGTC
Contig 37 (300 bp)
ACCTGATCAGGCTCTGCACTGTGTTTCATCAGCGGAGCGGAGATTTTGAC
CGCCCCATGCATAACGGAAAGGCGTGGSTAAACCCCGGGCGGTTCCCT
TATCAGATGACCTTCGAATATTCGGCAGGTGCAGTTTGTATTATCCAG
AAAGGCGTTGAGCGCGTATGAATATAATTTCTGTGGGATTTGAAGCATCT
TTTCCCTCCTTCGGTGAATGCGCTGAAACGGCTTATTCAGCGGTTCA
GGGTACGCTGATAATTTGCAATTTAAATACCATTATTGGETACTTTTT
Contig 38 (450 bp)
ATCCTTTTGGGCTCTGGCAATTACGCAATAAGAAGGCCCCCATGCGATT
AAAGTCACCGGCCACTGTCTGCTAATCATGGAGAAATTGTCCATCAGTC
GGGTCTCCATGGCAGGGGATTTGCTCTGCTTCTCTGTTGGATGTTAGCG
AAACATTGCCAGTGGTCAATTTAGTGCAAGTGCTACCGGAATATTACAG
CAAGCAACGCTCTGCTCCGTTTATGTTCAAGGCTGCGCAGCTCAGCGAA
AGTGGCGATACCGGTAGAGTTTACGCCAGTATTTGCGGAGCACTACC
GGATGTTTCACTGTGCTGCTGATTTATGATTCAATTTATCGGTTGA
TATCAGTTTAAACCTGATTTTCTCTTTCTAAGCCGCTACAGATTTGCT
AGCATATTCACCTTTAATCGCGCATGATCTAAGATAATTGAAGAGGTTA
Contig 39 (450 bp)
AATGTACTGGCAAAAGCCAATGGCGAAGCGTGGGGAACGTTACATGCTC
TGCTGGCGGATATTAAAGTACAGGTGAGGTGCAGATGGCGATGAACCGC
GGCATCTATGATGAAGCTATGCGCGCTCGCTTTGTACATCGAAACGG
TCAGCAGAAGGTGGCGTTAAATCTCCTTCAGGTGAAGGGAATTTCTTA
TCCGCTCTGCGCGCGTGTTTTATGTCGCGGGAGATAAAGTCGGCATCGTT
CGTCTGATGCTTCAAAACAGTAAAGAGATTCAAGTTTGGCGTGCAGTC
AGGGCCAATGTTGATGGAAAACGGTGAATTAATCCGCGTATTATCCCA
ACGTCCGCTCAAGCAAAATTCGTAACCGTGGTTGGGATTAATAAACATGG
GAACGCGTGTTTTGTGAGCCAGCAGGCAACAAATTTTATGATTTTG
Contig 40 (400 bp)
GACATTAATCATTTCAAAATCAAAGCCCGGTTTTCATCGCCGTTTGG
TGGCGTGGCACTGAACGCAATCGTTACGAGTGTAATAGTAATGCGCATG
ATTCTGATTTCCCTTTAAATGAAGATACGGCGCGATGATACCGCTCGGG
TTGCTCTCTGTTGATACAGAGATACTAGATGTAGTTGAAAAAGATTCA
ACCACACAATATATAGCCAGTAGGGGTGAAATTAACCTGGATATGAGC
GTGACGGGTAGGGGATTTTGTGATTACACGCAAAAAGAAACCCCG
AAGACAGGCTTCGGGGTCAAAGACCGGTATTTATTATCATTTTGCATA
CGATTTGCGCATGCTTAACAGTGGCGCGATTAAATATCTACCGCAGCTG
Contig 41 (500 bp)
GCAAAATCAGCTCCGCGACCTGGCSTTCTCGCTGGGCCATATTGGCAAAG
GAGCTGGATTGCGGTGCCTGCAAGTGCCTGAATAATGCCATTGTCTCTG
TACCGGAAGAAACCTTTCCGAATGAACACCCACAGCAGCAGCTAAGCA
GCAGCGTGCTGAGTGCCACGCTTAAGGTCAGCCACGGATGATTCAAGCACT
TTCGCGAGTCCACGACCATAGCGCGGATATCTCTGTCGAACATTTTTC
CGAGGACCGGAGAAAGCGGTTCTGTTACGCAACGACTCTGGCTGAGCA
TCCGCGCGCATCATCGGTGTCAGGCTCAGCGACACCACCGCTGAGATC

SUBSTITUTE SHEET (RULE 26)

FIGURE 6, CONTD.

AAAATCGCTACCGCCAGGGTAATAGCAAATTCGCGGAACAGTCGCCCCGAC
GATATCGCCCATAAACAGCAGTGGGATCAACACCGCAATCAGTGAGAAGG
TCAGCGAGATAATGGTAAAGCCGATTTCACCTGCGCCCTTGAGCGCCGCC
Contig 42 (400 bp)
AGCTATCTACGGCAAAAGGCACGGTAGTCAATTCGTTGTTAAATACATC
AAGCGTTTGGCGCCGAAATACCATCTGCCAGATGCCATTTCAATTCGTAG
CGCACTGCATAACGGCTACCGGATGCAGTACGTCAAACCCGAACCTGGGGC
CGGAAGGATTAGCTTTTCTGCAATACACCGCGCGCACCCTGGTGTGGC
GAAAGGCGCGATGCTGACTCACCGCAATATGCTGSCGAACCTGGAAACAGG
TTAACGCGACCTATGGTCCGCTGTTCATCCGGGCAAGAGCTGGTGGTG
ACGGCGCTGCCGCTGTATCACATTTTTCGCCCTGACCAATTAAGCTGCTGCT
GTTTATCAACTGGTGGGCGAAGCTGCTTATCACTAACCCCGCGGATA
Contig 43 (450 bp)
GATTAGCGCCAGATGCTCGCCATCGAAAAGTTGAATCAACCCAGCTGCG
GGTAATAAGTGCAGTACGAACAAATTCAGTATCCAGGGCTATCGCCGGA
AAGGCACGGACGGCTTCACACAAAGAGCCAGCGCATCGTCCGTGGTAAT
CATTTGGTAATTCAAATTTGTTTCTCTTTAGTGGCGCTCAAAAAAACGC
CGGATTAAACCGCGCTCTGACGACTGACTTAACGCTCAGGCTTTATTTGCC
ACTTTGGCGCGCGCTTCGTCACGTAAATTCGTCGCAAAATTTTTCGGAC
GTTAGATTTTCGTTAACTCATCACGAACTCCACCAGCTTCGCTACTTTGT
ATCCCGTGAGCTGACGGCGGCAAAAGTCAACAGTGACTCTTCGTTAAGC
GATGGATCTTTTTCACACGAAGATTTTCACCGCTTCACCAGCTGGAGCC
Contig 44 (750 bp)
GAGCAGCCCGCTGATGACAGGCATSCGCCCGCGTGGCTCTCTCTCTCT
GGTGCCTGAGTCACAGGATGCGCGCGTGGCGCGGTGGTGGAGCGGT
CCTGGAGGGCTCGGAGGGAGGATGCGCTCAAGCTGGCTCCCGTGGGGC
TGGCCCGGAGTAGCTTCGCTGAGGGCACCGTGTCTGCTCCAGAGCCCGC
TCCCGGCTGCGCTGCTTCCCTTCCCTGCGCTTCCCGCGGAGCGCC
TGGATCCCGATGGGAGGGCGCCCTGGGGAGAGGGGACAGGGAGGGGGCC
AGAGCTCTGAGGCCACACAGCTTGGCCAGGACCTTCGTGGGAAGAAGAG
GTGGGCCCCAAGGCACCTAGAGAGAGGGAGGCTCTGCTGGCTGGGGGC
CTTCCAGGGCGGGCTTCCAGGCAGGGCCAGTGTCTGGGGCTGGAGGGA
GTCCCTGCTGCTGGGGGCGGCGAGGACCTGGGGCTCTGGGAAGAG
AGCGGGAGGAGACTGGAGCCAACTGGGGGACAGAGGAGGGTCCAAACC
CAGCGGTGGTGTGGGGGTGCTGGTGGTGGAGGCGCTGAGAGGCTGTGCT
GGGGGCGAGAGCGGTGCTGGGAGGGGAGAGGGTCCCGAGGGCTCATG
GGCCCTTCGAGCAGTGGCAGTTGGGTGGGTGGCTGTCTTAGGGCTGT
ACCAAGTGGGTGCTGGAGAAAGAGGCTTACCCCTAGTCTTTGCTGCA
Contig 45 (300 bp)
TGGGGACCCCACTCCAGCCCACTGAGTACGCGCCCTCTGTGTCCCA
CCGCCAACCTGCCTCACACCAGAGGGGCTGTGGCCACACCTTGTCCACA
GCCTGTCCCTGAGACCACAGCCCGGGCTCAGCCCTCTCTCACCCCT
GGACCGAGGAGAAGCCCACTGGGTGAGCTCTTGGAGCTAAACTTCC
AGGAAGGTTCTGTGCTTGGGTCTTAGAGCATGGTGGGGAGGGGGATG
CTGGTGGGGCGCAAGCCCTCCCACTTTCGCACTCGACCCGGTGGGNG
Contig 46 (300 bp)
CGGGCTAGAAGCCACGAGAGCCCAAGGCCCCGCGCCGAGCTCTCTCTGC
AGGGATTGCGCAGCCCTGGGGCCACAGGGCTGAGCAGACCTTGGGGTTC
CGGTGTGACTCCAGCCAGGGTCCCTACTGTGTAGGCACAGGGCAGAGTC
AGCCCTGGGACCATGGCACAAGCTGCTCCCGCTGAGCCGGGCCCCCGC
CCAGGCTGGGCCCCCTCAGTGCACTGTCCCAAGCCAGCTGCTCTCCAC
CTCCACTTCTCCATCCAGGTCTGCCCCACGGCTTTGCTCAGGCCACG
Contig 47 (500 bp)
TTGACTGGCACTAGCAGGCTCTGTACCCGGGATCTGGGCTCGGGAGA
AGGGAGACCCCAACCCGCGAGGCCGAGGGCGCTGTACACCATGACTCT
CAGCCTTCCCAACCCGACGGACAAGATGACCTCTCCCAAGCCCACT
CAACCAGGACCGCACACCCCTGAGTCTGCGAGTGGGGGCGGCTCAGGG
GCCCCGAGTCCCAAGGAGTCTGCTGGCCCTGGGGGGAGGGGAAGCAGC
AGGGTGGTACCGGTCTCCCTGGTGGCAGGACCAAGCTCAGCCGCT
GGCTCCAGAGGGCAGCUGGACACCAACAGTCCGGGACCCACGTACC
TCAGCTGCTGCAAGTGGCCCTGCTGTACTGGTGGCAATGGGGCGGCTGG
CTGCTCCCATGGACAGTTCGCCACTCATCCAGCCGCTACCCCTTCC
GGGTCAAGTGTCCGGCGGGCACCCGCTGCCAGCCCTGGGCTCTCTC
Contig 48 (500 bp)

FIGURE 6, CONTD.

GGGGTTGCCGAGGCTGCTGTGTAGTTCGACAGCAGCTTGGATCTGGC
GTGGCTGTGGCTGTGGCTGTGGCTGTGGCATAGGTCAGCCACTGCGACTC
CGATTTGACCCCGAGCCCGCACTCCACATGSCACAGGTGCAGCAGGG
AAAAATAAATAAATGAAATAAAATAGGTGAAGACAGTGGATTTCATCTCT
TGGGGTTGCGGTAAGCTCTACACAATAGGGAGTTTACCATTTCACCTGTT
TCAAGTGGCACTGAGTCAGCTCACAGTCCTGAGGGCCACAGATGCCGTC
TGCTTGGGAGATTGTTCTCTCACCACACTGCCCTCTGTCCCCACTAAA
TACTCACTGCCCTCCCGTCCCAAGGGCCCTGCCACCCCTCTGCTTCC
TGTCTCTGAACCTTCTGGCCACCAGCGACCTGTGGTGACCTCACTCTTC
GGCCCATTTGTGCGACACCCCACTGGCTCTCCCCGGCATGGGCAGAN
Contig 49 (600 bp)
GGCATATTTGGGGGCATATTTGGGGGGAGATCCCAACAAGGCATTGGG
GTTTGTGGTTTGAATGCCCGGCCCGATGGAGGGGGCGGGGAAGAA
TCTAAGCCTTACTTGGGGAGGGTTGGGCCCCGGGGCCCGGGCCGAAAT
GCCCCAAGACAGAAGGTGTACAAAATTTCTCAAAGGGTGACCCCTAAT
GAAACGGGTCCCGTTGAAAGAGGTACCCAGGGTGGATTGGTGGCACC
CAGAAATTTACGACATTTTGGCTCTCTTCCAATGGCCGACGCTGGGGAT
AGGCGCCCCCGTGGACGGCGGGCTCTCGGGTGGGACGGCGGTGAGGGGT
CGGTGACGCTTGGCTCTCTGACCGCTTCCAGCTCCTTGGGAGCGTGG
AGCGCGGGGGCGCGCAGGACGGCCGCGCAGGCCCTTGGCAGGCGTTGG
GCGGACTCTTCCAGGTGTATAGCGGAAGAACTTCCACGCGGGTATCT
GGGGAAGTTGTCTGAGAGGGGAAGGGCCCGTCAGGGGGGGGCTGGCCC
CCAGCCCTCTGCCAGAACAACTTTGCGGGGTCTCTGCTGCC
Contig 50 (179 bp)
ATCTTCATATTCATGCAGAACACCTCTCTGCTTTCTATCTTGGGGAA
AAGGACGATGTCACTTATGCAATAAGGCCACTTGTGGCCGGGGCTTGA
CATTATTCCTTCTGTCTGGCTCTGCACCGTATTGAACCTGAGTTAATGG
GCAAAATTTGATGAAGTAACCTGCCACC
Contig 51 (500 bp)
CTCGGCTCTTTCAGGGGGCTTGGGAGCCATAGAATGCTATGGAGCA
AGAGAGTGTATGGTACAGACGACTTTGGGGGAAGGTCTGGGAGAACAGGG
GTGACTGCCCACTGTGATAAAGAGTGGCGCTTCTTGAAGATAACACGGT
GGGACGCGAGCTGACCTGTGAGGTGAGAAAGGCTCTTCCCGCGGCC
AGTACGTGGCTCTGGGCTCCCGACACGAGAAAGCCACCTCCACGGCTG
CCTCCAGGGGGCTTCTCTCTTACACCGCGGGCCATGCCAGGTGTC
AGGTGCCATCAGAGGGTGTCTCAAGAGAAGCTCTGGGCTGGGGTTGTCCA
GGTCCCGGAAGCCCGTGTCCAGGGGCCACTGAGGAAGCGTGGGGCGCA
CAGAGACTGTCCCTCGGTGCTCAGAGAGGGTCCCGTCCCAAGGCAACGA
CGCCCAAGGCGCAGGTGTGTCAGAGGTCTTGGGAGGGAGGTGGCGCGCA
Contig 52 (900 bp)
TGTGTTGCACCTGTGCTGCCCTGTCGACTCTAGAGGATCAATACTCTTA
CATAATTAAGAGAACAAATGGAATTAATAAATGATGGGACATATTT
CTATTATCCCGATTACAGACAGCTTGGAAATGGAACATAAGTTATCG
GATAATCTACTGTTGACTATTTGTGCCGTTATTTCTGGTGACAGAGGCTG
GGAAGATATAGAGGATTTTGGGGAACACATCCCGATTTTGAAGCAAT
ATGGTGATTTTGAATGGTATTCCTGTTACAGACACCTTGGCAGAGTT
GTATCTGTATCAGTCTTCAAAATTTACAGAGTGTCTTATTAACCTGGAT
GCGTGACTGCCATTCTTCAATGATAAAGACGTCATTCGAATTTGATGGAA
AAACGCTCCCGCATTTATGATAAGAGTCGCCGAGGGGAGCGATTTCAT
GTCATTAGTGGCTTCTCAACAAATGCACAGTCTGGTCATCGGACAGATCAA
GACGGATGAGAAATCTAATGAGATTACAGCTATCCAGAACTTCTTAACA
TGCTGGATATTAAGGAAAAATCATCAAACTGATGCGATGGCTTGGCAG
AAAGATATTGCAGAGAAGATACAAAACAGGGAGGTGATTATTTATTCGC
TGTAAGGAAACAGGGGGCGCTAAATAAGCCCTTGGGAAAAATTTTC
CGCTGAAAGAATTAATAATCCAGCGCATGACAGTTACGCAATGAGTGAA
AAGAGTCACGGCAGAGAAGAAATCCGTCTTCATATTTGCGATGTCCC
TGATGAACCTTATGATTTACGTTTGAATAGAAAGGGTGAAGAAATTTAT
GCGTGGCAGTCTCTTTTGGTCCATAATAGCAGAACAAAGAAAGAGCTC
Contig 53 (450 bp)
CCAGCCACAGCTGGACCTTCCCGAGAGGGGCTGCCCTCTTTCCCGC
CCAGACGCCCCCAGCAATCTGTGGCCAGAGGGAGTGATACCGAAGATG
GCCACATGGGGGCGCCAGCCACAGGGAACCCAGGAAGGCGCTGGACCG
TCAGGAGTCAGGGCTGCTGTGCACCATGTGGCTGGGGACTTTCCACAG
CCTGTTGGAGATGGCCGGGCACACCGCTGCCCTGGGGGAACGTGCACAG

FIGURE 6, CONTD.

GGTGGTACATGTGGCCGGAGCCCAGGGCACAGGGTGAGGGGAGAAGGGAG
CATGGCGGTGCAGACTCGGAGCCCGCGCTGAGGTGCTGGGTCTCAGGA
CACGCTCTGGGAGTGGAGGACCCCATCCACGCCCCACCCAGTGTGTGC
CCGCTGCTCCCCGGAAACCTCACAGACACGAGGGCACACCCAGCCCC
Contig 54 (1133 bp)

ATGGCGCTCATTAGAATTCCACCTCGGTACCTTGGGATCTTTTGACCCCT
ACCTCACGCCATCTACAACATTTACCTCCGAATGAATGAGAGACACCAAA
AGCAAAATTCATAGAAGAGAAAAAGGTAACCTGGACTTTAAAAATGTAA
ACTTCTGCTCTTTAAAGGCAGTGTCTAATGAAGTCAAAATACAAACCACA
CACCATTAAGAAAAATCTTGCAATCTTGTCTGACAAAGACTAGTGTICA
GAACATACGACGATCAGGGAGAGGAAACAGCAATCCTATAAACTGGA
CAAAGAATTGGGGGAAAAAAAACCCACTTGGCCAAGAAGTTGGTAAATA
AGGCCATGAAAAATGCTCAACATCATGAGTCATPAGAAAAATGCAAAAT
AAAATTATAATGAGATACTACTACACAGCTATTGAAATGGATAAAAAATG
TTTTAAAACTGATTATACCCAGGTTTGGCAAGACATGAGAAACGAGAT
TTTACACACGATTGGTGGAAAAACAGAAATGGTCCACCCACTTTGGAAA
AGAGCTGGGCACTTCCCTCAAAGTTAAACATACATCCAGGACCTCACAC
AGGCTTTCCACCACAGGTGTTTATCCAGACATGAAAGCGCTCATCCA
CACAAAGACTCGTAAATGAAGGTTTATAGCACCGTTTGGGGCCGAAGT
AGAAAAACCAATGACCTTTAACCAGAGATATCTAAACAAATATCCAT
TCACATTAATCACCCATAAGAAGAACGGGCTATGGGGACGGGAACCGTA
TTGAAGAGGGTCAAAATACATACGACGATCAAAGAAGCCTGCCCCAAGG
ACACACACTGCAGGGTTCCATGGACTGAAACTCGAGAAGGTGAAAACTCG
CCAGCAGTGACACAGACGAGTCCGACATCAACCTGATGTGGAGGAAAGT
GAAACCTCGTGCGTTGTTGGCAGGACTATAAAGTGGAGCAGCCCTACGG
ACAACAGTAGCCCGGGCTCCTCTCCTCATCTCCTGGGGAGCCTGAGCC
TTGAGACGCTGGGGCAAGTGACGGCATGCTGCTCAGCTGGGGCCCCGG
TGAAACACGTGGCAGCTGGGGAAAGATCGTA

Contig 55 (735 bp)

TACTGCCTGTCTCTATGGACTTGACTCCTCTCGGGACTTCATGCGAGGGA
TCTTACAGAAATTTGTCTTTTGCACTCTGGCTTGTTCAGTGAAGCATCGTG
TCCCCAAGGTCCATCCATGTTGCAGCCTGTGTGAGGATTTCCCTCTCTTT
CAAGGCTGAATAGTACTCCACTCTGCGGATGGACCAGTTTGTATTATCC
ATACTAGTAAATCCATACTAATACTTGTTCAGTGAAGCCACAGCTTAT
GCTACCTTCCGTGGGCTCCTCCCTGCCCTGTCTACGCTTCTGTCTATA
GCCCATCCCCCTCTCATCCAGGCCACGCTCCTGTCCCTGGACACTGT
CCAGAAGCCAACCTCCCTCTGACTGTGCTCTCGGCTGACGGAGGACAAG
GCAGGCTCAGGGGTCCACGGGCTGGGGCCCCAGGCTCCCATGGTGTGT
GCCCTTCTCTGATTCCAGAAGTACAGTGGCAGCACCAGCTTCCAGCTGC
CCCACCTTCTGTCCGAGGCTGCTCGGGTGGGGGACAGTGGGCAGTGATG
TCACCTGCTGTAACCAACCTACCGTCTGCTCATCCTGTCCAGGAGGTAC
GGTGACCTTGGCAACATTCTGAACAACACACCTCCCTCTGCTTAGAG
GCCGGGGGCTCCCCGGGTGACTGGGGGACAGGCTGACCCAGCCTGTG
TCTCTTCTCTGAAGGACATGATAAGTACTGCAACA

Contig 56 (500 bp)

AGGAAGAACAGGAAACAACGGCTTGAGCAGAAGAAACGGGTGTCTGGCA
GGGGCACTGCAACAGGTCCACCGGGTGTGCGCGCTGCGGCTGCGGC
CAGAGGGGGCAGCTCCGCCCTCGGGCCGCGCCTGCGCTTGTGTGGC
TGGCGGCTGGGCTCTGCTTGGCTGGGTACAGCTGGGTGACCCGAGGC
TGTGGTGGGTGCCGCGGGTCAAGCAGCCCGGCCCCACCGGCGGCTCTC
GCCGCTGGCCGGGACGCTCCTGCACTGAGGAGTCCGCTGACGG
GCTGATTGGTCCACAGCCTCAGATGCAAAACAGCCCCAGTGCCTGGAGC
CAGCCAGCCCGGACACCTGGTGGAGGACGGAAGGCAGCAGCTGGAGA
GCCGCGCGGATGATGCTGCGGGGAAACCGGCTCCCGCGGGGGCGGCC
TGGCTCTGGCCAGGCTTGCTTTGAATGCTGACGTGAGCGGTGGCCCTATA
Contig 57 (500 bp)

TGGCGTTGCAGTGGCTCTGGCGGAGGCGGGCTACAGCTCCGATTGGA
CCCCTAGGCTGGGAACCTCCATAAGCTGTGGGTGACGCCCTAAAAAGCAA
AAAACCCCAACATATATATATATATATATATATATATATATATATAT
CATAAAAATAGAAATTTACCTTCTTAATAATTTTCACTGCACAAATTCAGTGG
CACTAAGCACATTCTGCGGGCGGTGTCACCTGCTCCAGAACTTCCATCT
ACCCAAACGGACTCTCCGCCCATGGAACACGCCCCCTGCCCTCCCCCG
GCCCTGCCCGCCAGCTCCTCCCTGTGTCTGTGGATCCGGCTCCTCCAGG

SUBSTITUTE SHEET (RULE 26)

FIGURE 6, CONTD.

GACCCCCGTGCGTGGGCTCACAGAGTGTGTGTCCTCTGTGACCGATCGTC
GTGTCCCCGAGGCCCGTTCTGTGGCAGCTGCGTTATGACCGACTACCTTC
GAATGCTCAGTGACTGCCGTGCATTGGACACGCAGTCCGCTACCTTTTC
Contig 58 (550 bp)
TGCTTTCTGTGCCCCCTCCAGCTTGGGACCCAGCAGGGCAAGGGGTGT
ATAGGGCTTAAGGAGGCAGGGGGCGTCTCCTCCCGTGGCTGCCAGAGC
ACCCCCAGCCCCGCTGCCCTCGTCCATCTCCAGCCTGTCTTTCTGT
GCCCTCCCTGTCCCGGGCGGGCCGACACTGGCTTCCACCTCCCCACCCA
ACTGGCGGGCCCGTCTTCTGCTGAGGCACCCGAGGTCCCGCTGCTG
GGGACCACTGGCAGGTGGGTCCCACTGCTTTCTCAGCSTGGGCTTTGGA
GGGGGGATCTGCACATACCATCCCTTCAGGCCCGCTGGGAGCCTGGGGA
CCATCCCGCACCCCTGTGGGCAGGCCAGAGGACTGCCAGGAAGAGACCC
AGGGGACCAAGCAGTCCAGGCCTCTCAGCTTCAGGCCAGGGGAGCCCA
CCCCAGGTGGCAGGTGAAGCAGGCCCCCAACCCACAAACTGCCCGCA
GGGAAGTAGGACGACAGGAGGGGAGGCCAGGCCCGGCCCTCTTG
Contig 59 (800 bp)
T'GAGGAGCGCAGGCCAGGCCTGAGTGTGCCAGCTTACACCCCTGGCAG
CTTCGTCCTCTGGCCCTAACCCCTATCTTACCCAGCAGCAGGGGCTG
CCCCGGTGGGGCTCTGCTGAGCGTCTGACTGGGGTTTGGAGTCACTCTGC
TCCAGGCTCAGCCCCATCCCAAGGCTGCCCTGCAGCACTGTGCCAC
CCCCATAGCGCCCGCAGACCTTCGCCCTCCAGCCTGGATGTACCCACGGA
CCCTGAAAAGTGGCGCTCAGCAGGTGCCCTGGCTGGAGTCCCGCTGACTT
GGGCTCGCCAGGCTGCCCTGGAGGGCTGTGGGGCACAGCCTGCCCA
GGGGCCCGCTGGGCACTGGCTCTGAGCTCACGACAGGCAGGCCCTCTCT
TCTTGGCGGGCCACACCTGCCCTGGGGTTTGGGCTCAAGCCGGGCAGC
CCCATGTAGGCGGGGGCAACCAAGTAATTACAGCTTGGCAGCCGCT
CCCCAGACCCCGCAGCCCGGAGGGGCCACCCAGGCTGTGCCACCAAGA
CTTGGCATCCAGGCCCAAGCAGGTCAAGGGCAGCTGTACAGATTCTT
TTAAGTTGAGACAGAATCGACACATGACAAGTTCCTGGTTTAGGTACTT
CGCTGCCGGGGCCGCACTCAGTTTAGTGACCCAGCACACCCACACAGG
TACAA'TTGTCTTCTCAAAAGAGGCCCTGAGAGAGCGCTGTCTTGGCT
CAGGGGTAATGAGCCCAATGGGTATCCATGAGGTTGCCGGTTCATCCCC
GGCTCGCCGCTTGGTTA
Contig 60 (500 bp)
GGCTCAGGAAGCCAGGGCCAGCCTGTGGGGCGACGGGAACCATGGGGGT
CTGTCTTCCCGCTCTCTCAAGCCACCGGCTGTGTGCCACCTCCGAC
TCTGCAGCCAGCATGCCGGCTAGAGCCCTGTGCACCCAGCTGGTGGCT
CTGGCTAAGGGCAGTGTCTGGCTGTGGACGGTGTCCCTCCCGCAGAGCC
CAAGGGTCCCATCTGCCAGGCTGGTGGCTGAAGTCTTGCCTGTGTGGTCC
TTGCAAAACCCCGCTCTCTGCCCTTGAAGCGTCAAGGAGACGGGG
GCTGGCGGATGCCCTCGGGCACAGCCCGCCGCGTGGCGCCCTGTGAG
GAGGGGGCTCCGACSTGCCCTGACGGCCCTGGCGGGCGAGAGGGTGAG
GCCACCTCTGGCCACGTCCACCCAGCTGCCACGCCGCTAGCCAGTGCC
CGGGGCCAAGTCAAGCAGAGCCAGCTTCCGACAGCAGAGGCTGTAGGC
Contig 61 (700 bp)
GATGAGGAAGCCGCTGCTGCTGCTGCTCTTCTTGGCCTTGGCCTCGT
GCTGCTATGCTGCTTACCGCCCGAGTGAGACTCTGTGGCGGGGAGCTG
GTGGAACCCCTCCAGTTTGTCTCGGGGACCGCGCTTCTACTCAGTAA
GTAGCTCAGCGGGGCACGGGGCGGGGGGACACAGCAGGTGCTCCATCG
GTGCTGCCCGGTACCTGTGCGGGTCTTCCGGATGGATGGTGTGGGGGA
CGGGGGCGGGGGCGGCCAAGGGAGGACCTCTCTCCGAGGGTCTGAGA
CTTCAGACCGGGGGCGCCCTGCCCTGCGCATTTGATTGGCACCTGCCATG
TGCCTGGCTGGGGCTCACACCCCTGACGTTCTTGCAGCGTGACTCGAAA
CGGGAACCGAAGGGACCCCTGCCACCGGGTGGGGAGGCAGCCGTGAGT
GGCAGCGTGGAGGGGTTCCTTCCGGCGGGGTGGCCAGGCAGGCCCA
CAGGATGACAGCCTGTCCCTCTGCTCTCTTGACTGCCCCACAGCCA
GGCTGCAGGCACTGACATTCACCATGGTATTGTGGTGCCTTGACGTCT
TGGCAGTGGGCATTGGGTTCATGGACTGTTGGATTGAAAAGTGGGAATA
AGATGGGGTTTGA AAAACCAATTAAGAAATAAAGGGCGCCCTGTGGGC
Contig 62 (300 bp)
TTTGA AAAATTTTCACTCAGTGACAGATTTCGCATCTATTCGGCATTGAGG
CTCTCTGTCTCACCTTGCTTGTGCGGATCTTCTATAACCAACACAG
TGACGTTTTCAAGTACTTTATTGAATAATAAGAAAAAGTGACACAAAT
CATGTAGTTAACTTTCTGTGCTCTTGGCAGTTTGAAGGGACCTCTTTT

FIGURE 6, CONTD.

TTTCCTTTTATAGGGCTTCCGCSACCGAAGTTCCCGGGCTAGGGGTTGAGT
CAGAGCTGCAGCTGCTGGCTACAGCACAGCTCTTGGCGCGATGGATCC
Contig 63 (450 bp)
TCCTGGGCCACAGGCTGCAGCAGCTCACCTGGGGGCTGGGGTCTCGCTCT
GCGGATGGACCCATGAAGGCCGAGCCAGGTGGGGGCCGAGACGGCAGGG
CAAAGGGTCTGCACACACAGCGTCCCCCGACCCGGCTTCTTGGGTTCT
TGGGGGTTGGCGAGGCTTCTCTCAGTCTGGGTTCTCTGGGAACTTTCA
AGAACTGGGAAGTCTTCCAGAAAGTTGGGGTGAGGGGAGGTACCCCCAAA
GTGCTGCTCCTCTCCCCATCCCCACCCCGCTGTCCATCGGCGAGACCCC
GGACCGCGTCTCCTGCGGAGGTGTGGGGTCCCCCCTCTGCCGCGCAG
CCTGGGCAAGGGTGAGCGCCCCCTGCTCTGCACTCGGGACTCAGCCTGGG
GAAGGCGGGCCCCAGGAGGTCTGGCTGGACGGCAGTGACCTTCCACCG
Contig 64 (500 bp)

TGTGCATCAACCCAGTGCCACGGGGGTGACCTCGGGCCGGTCAGCC
GCCCGGCTCTCCACGGAAACCGGGCTTGGCTGAGGCAGAAGGACCCAG
GACTCCATCCCTGCCCGGACTCTCCGGAGGGTGCGGTCTGCACAGAGA
CCCTCTGGGGTGAGGCCGGTGGGGCTGGGTTGAGATGGGATGGTCAG
GGCGGCCCCCGGGGCTGCAGGAGGCTGGGTGAAGGAGGGGCCAGCT
CAGACGCCCCCAAACCTAGCTTGGGAGAGCTGCAGCCCCCGGCTCAAT
CGCGACAGCCTGCCACAGAAGGCTTCAAATGAGAGACAAATATTGGG
CTGAAGACTATACCCAGCCAGCTCTCTTGGGAGCCCAAGCTGCTCCA
GGCCCTCATTTGGGTATTAAATTCGTTTCTTTAGAGATTGCGATGCTTA
TCAATGGCCACTGGGGGCTGGGCTGGATGCGCTCCAGGCTTTGTATG
Contig 65 (661 bp)
TCCCACGACCTGCCCTCCAGGGCCACATCTGGCGACACCGTCCCAAGAG
TTGGACCGGCTGCTGTGGCCACAGCCTCAGGCTTGTCTGGCGGCCAG
GCCGGCTCCAGGCTCCAAGGAGCTCTGCTGCCCTCCGGAACCCAGCA
CCCCGGGCGGCTTCCCCACAGACCTGTTTTCCAGGTCAAGGTACAG
CTAATTTGGGCTTAAACTGGACAAGGAGGCTTATCTGGAGCAGGCTCCU
GGCCCTTTGGCTCTGCCCTGCTGGGAGGCTTCCCAGAGGCTGTGTGT
TGGCGCTGACCGTGACGCTGAGCTTGAACCCGGATAAGGAGGACCCC
ACCTGGGCTGGAGCCAGAGAGCCCTGTTCCCGAGCTCCGCAAGGTTCTC
ACAGTCCCGCCCTGCCCTGGGGACCTGGAUCTCCCGAGCAGGTGAAG
GTCCAGATGCCCTGACTAGAGGCTCTCCGCTCTCAGACATGCTCCCT
TCCCGCACCGAGGACGAGACCTCAGCAGCCCTGCTGGGCTGGGGTGGG
ACCCCAAGGCTCTCTGAGTGTGTTCTAATGGGGAGGCTGGGGCTCAA
CAGTGGGGTGGCACTTGGAGGGAGGCTCCCCACAGCTGCCCAAGATG
GGCCCTGGACT

Contig 66 (500 bp)
TTTGTGGATGAATGAAATCATGAGAAAGTGATTGGACCGCCCGTTCCT
CCAGCTGCTTCCAGCTGCTTTGTAAAGATGACCTCTCAGCTTCTCAGAG
GCTTGGCGGCGCCGAGGTGGCAGTCAGTGAGATGCCATGCTTTGTTGGC
ACGTGGGAGGCCCCGTGTCCACGGGCTGGGTGCTCTTGTGTCTAATCAGG
GTGAGGGGAGCAGCAGGTGCAGGGCAUATGTGGGGCCGGGCGGATGTC
TGGGGAGGGCGGGAGGAGGGGGTGTGCGGAGGCGCTTGTGGGGTGCAGG
GGACAGACCCAGCGAGACCCCTCCCTGGCCAGGCACAGGACAGGTGATG
GGGGCCCGCTCCGGGGGCTGTGACAGAAGCTCTCAGAGGAGGCCCCC
CACGCTCTCTGGACCATCAAGGGACCGGGGCGCTGGGCTGGGGGTAC
ACCCAGCTGGCGGCAAGCCCGGGTGGGTGGGAGGCCCGGCGAGTTAC
Contig 67 (550 bp)

GGSCAGGAGGGCCCCGGGGCTGGTGCAGCGGTGGAGGTGGTGCAGGAGG
GTGTAGGCGAGGCTCACTGAGCGTGCAGGCTGGCTGTGCCCTAGAGTG
GTAGACGCTGCCCCACCCTCCAGTGTGCTCTGTTACCTGTGCTGG
CTCACAGGTGTGAAACTGAGACTCGGGTGTTCATGAGCTTCCAGGATG
AGAATCAGCAGGCTTCCAGGCGAGGCTGTGTCCGGGGCTCTGGGCTCTT
ACCAAGGAGGGGACACCCAGGGACAGCCCTGCTTGGGGTGTCTGGGCTGG
CCAGGCTGGGTGGTCTCTGTGGCTGGCAGCCCTTGGCAGTACCCCC
TTACCTCAACTGCCCTCAGCTGAGACACGACCTCCCTGCAGAGCCCTG
TCCACCCAGACATCACCTCGCTCTCTCAGGAAGCTTCCAGGGCTGCT
CGCCCTGGTCTCAGCAGGAGACAGAGAGAGGGTGGGCCAGGAGCAGA
GGCAGGCGAGGAGGGGAAGCCCGGGGCTCACTACCCCCGGGGCC
Contig 68 (500 bp)

TTTGCATTACGCTCGTACCGGGATCCTTCCCGGGGCTCTGGGGTGGG

FIGURE 6, CONTD.

GGAATGGGGTCAGAGGCAGCTGTCTATCTGCCTGTCTACCTGCTCTCAC
AGGCTGGCCCTGGAGCCCTGGCCCTCCTTAGGGGCACATCAGGTTTTGG
GGGAGGCCAGCCACCGTCCACCTCCAAGACCACAGCTGGGAGCCCTGC
CCCCAAGCCTAGACCTAGTGGGGCTCCTGCCAGCCAGGCCCCACCTTC
ATGCTGCCACCCACCAAGGTGGACAGTGCAGCCAGGACATCCAGCTTCT
GGAGCTGCCCGAGGCTCAGCACAGGCTGGTACCTAGGGAGCAGGTCACC
CAGGGCCGCTGGCGAGGCTGCGGGGACGGGGGTAGGGTGGGCAGCAA
AAGAACCTCTGAGCTGGGCGGGGCGGGTCTGGTGAAGGCCGGGGCCG
GGCTGTGTGCGTGGCCCTGAGCCGTGCAGACGACAGCCCTGGGTGGGT
Contig 69 (550 bp)
TGTGCTGCTGTGGCTGTGGTGTAGGCCCGCAGCTGCAGCTCTGATTCCGA
CTCCTAGCCTGCGAACCTCCATATGCTGCTCTAAAAAGACAAACATAAAA
TAAATGGGTGGCGCTGTATTTGAACACTCTGCCTCCTCCAGAGACGAG
GCCGAAACAGGCCCTCTCTGAAGGTCCACCTGCCAGGGAGGAGGAGGCA
GCCCCGTGGGGGCGAGACAGAAGCCCGATGTCCCAGACACACAGCACA
GGGACCGTGGCCCGGCTGCCAGCCCGCGGGGGAGGGCAAGGCCAGAG
ACTCCAGCAGCCACAGGACCTTGGTGGCCACAGGACACAAACACAGGT
GACGGTGGGTGAGGCTTGGCCCTTCCCCCTGGGCACAGCAGGACACA
CACAAGAGCCCGAGCTGCTGACCGCCAGCCCAAGGAGCCTGGATGAAGC
TGGACACCGAGAGTCCACACTGTGTGATTAGGCTGACGTGAAGTTAAGA
ACAAGCGGTGGCTCAGCGCTGAAGGCCAGAACAGGCCGGGAGGCGCAG
Contig 70 (1300 bp)
ATGTCAGGATAGTAACCTGGGGTGTGCAAGTGACAATGCCAGATCCCTTAA
CCACTGTGCCACAGGGAACCTCCTTGACCTAGAATCCTATACCCACTGCA
AATATATTTCAAAAAAGGTAAAGTCTGAGCAGAAAGCAAAATGGGAT
AATTCATTTCTGGAAGACCTTCTTCTTAAAGGAAGTTTTTTGGACGTGA
TGAAGGTAGAAATCGGAGGCACACAAAGAAAGAAAGAAAGAGCAC
TGGAAACGGAGCAAAATAAGGTAAAAATAAGTTTATCTCTTTCTCATTT
TTTAATTGCTCCAAAAGATAGCTGACCTCTAAAGTAAAAATAGTGA/AA
TGTAGCATATGTCTCTAGCGTAATTTAAAGTATAACTTTATAGCAATGATA
GCCCAATAAAGGAGGAATTGAGAATATACAGTTGCTGTGTTCCCATTTGT
GGCTCAGCAGTAATGAACCTGGCTAATATCCATGAGGATGCGAGTTCAAT
CCCTGGCCCTCACTCACTGGGTAAAGGATCCAGGGTTGCAGTGAGATGTG
ACGTATGTACAGACGTGGCTCGGATCTGGCATTTCTGTCACTCTGGCTG
TGGTGTAGGCCAGCATCTGCACCTCCGATTTGACCCCTAGCCTGGGAACC
ACCATATGCTGCTGGTGTGGCCCTAACAGACACAAATAAAATAAAATA
AAAGAGAGAGAGAATATACCATTTGAATTTCTCACATGACACAAAGAG
CAATGTGATATTATTTGGTATATGGTGATTGATTCAAGATGTATATCATA
ATATTGATTCAAGATGTATATATTCCTTTCTAAAAAGAGATTTATACA
ATAAGGCAAGAGTGAAAATAAAGTGGAAATGCTAAAGAAATAGTTAATCCAA
AAGAAAGCAGAAATGGGGAAAGACATATAACAGATGGAACAAATAAA
AAGAGCTAATGAGATTGTAATTTAATCCAAACATACAGATAATCCCAT
TAAATTTAAACACTCTCAACACATGATTAAAGAAATTTGCAAAATGAA
TAAACAAAGCAAGACCCCACTAGATGCAGACTATGAAAACCCACTTCAT
ATAAAGACATGGGTAGGTTTAGAGCAGATGATGGGCAACCATGTACAG
CAAAACATTTGTCAAAATAAAGCTGSGTGGCTGTATTCTATCTCAGACACA
GCAGACTTCAGAACAAAGAAACACTGCAAGGATGAAAGAGATACTGCATA
ATGATAAAGGGATCAATTTTCCAAGTGCAGGCTCCAAACAACAGAGGTTT
Contig 71 (500 bp)
ATGACCTCATCTGAATCGAGCTCGGTATCAGGGGATCTCTCAGCTGGGG
GGGAGGGCAATGGGGCATTTGTCTGAGGATGCCCGAGGCAGGCCCATTTG
GCTGGTTTGGTGCCCATGCCCCCCCCACACCCCGCAGTGCCCCCTGCTG
AGCCTGGGACCCCTCTGGGAGTTAGGATTTGGGGGTGGGAACAGGCTT
TGCAGTAATTTCCAGCCCCCAGGGCCCTTCCCTCCCGCCCTCAGGACCC
CAGCCCCGCCCCACACAGTCTCCACTGTGACAGCCTCACCCCTTGGGTCA
AGTCTGTCTCTCCGGCCCCCGCTGGGCAGTGGAGCCAGCTAGGTGAGA
GGCACAGGCCACTAGGGCGGTGGGCACTGCTGAGGACAGAGGGGCTGGG
TGGCCTTGGACGAGGCCAGCGACGCTGAGACAGTGAAGCAGGCTCCAGG
CTTTCCAGGGAGGGTCCCTGAATGTCCACTTCTTGTGACATCGGGTGAC
Contig 72 (550 bp)
AAGTCCATTAGGGAAGGGATTTGTGCAACACAGAGACAGGTGCAGGGCT
GGGCCAGCTGCTGGGCTGGGGCTCCTCAAGGCCGCCGTAAACCCCTCCC
TGCCAGCCGCTGCCGCCAAGGTCTGCTGTCCACCCGGCCGGGCTGCTG
TGTTCCCGCGTGTGCTCGGAACCCGACTCCCGTTCACCCCTGAGCAC

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FIGURE 6. CONTD.

TGCTTGGAGGCGGCTGCCAGGCGGGACGGGCCCTCAGGGCTGGGCTGG
CTCTTGGCCTGTGTTTCATTTCAGCAGGTCTTCTCAGTGGGGGGGGC
CTTGGGTGAAGCAGGCATGTGCACCACTGGGGCCCTGTCCCCAGTGGGCA
TCCTGGGCGCTTGTCTGGGCCCCAAACCCCAAGCCGTGTGCATCATACC
TTCACCTGAGCCCCAGCCGAACCCCGACATGTGCTGGGGGACCTGGG
CACAGGGGTGAGGGAGCAGTGGCCTTGGTGGAGGCCAGCCTTGGCACCT
GGGAGGGGGTGCATCTGGCATGCTCTGTGTAACCAAGCCAGGGCAGG
Contig 73 (950 bp)

GACGTGCAGTAGCCATGACCTCTACGGCCCCACTGACCAGCCCGTGTCC
TTGTCCCAGAGACCGACCCCTAAGCAATAGGATGCAGCAGAAGTGACAGAA
CGGCCTCCGCGATGAGGTCCGAGAGGCTCTGGCTCTGACTCAGGCCCT
CATCCCTCGCTCTCTGGAGCAGGGCCAGGTAGGGGCCCCCCAGAGACGC
CTTAGAGGAGGTGACGGGCGAGCCAGCCCCCCCCAGGGAAGGCTGGGGAC
ACCAAGGAACAGAACGGCACAGGCTCTGGCACAGTCTCCAGGAGCCCC
CTGGTGGCACAGAACTCTGACCGGCCAGTGGAGGGGCTGGGGCGGG
CTCGGGGAGGAGGACTGGGTGAGGCCGTCTGACTCTGGCTGAGCGCCG
CATACTGTCTGCTGCCACGATGCCGGGCCAGGCTTCCGCACGGACCC
AGGCTCACATTCGCCCTACATGCCACTGTGTGGGAGTTTGGGATGGTGTG
CCCGCTGGGCCCCGGGGTCAAGGCACGCTTCCAGAGGAGCGGGTTCCAG
AAGGCCCAGGTGGAGAGGCGATAGGAGGGCTCCAGGGGCTTCCAGGCC
ACCTGCCAGGACCCCTCTGGGGGGAAGGGAGCGGAGGAGACAGCCGGGT
CCCTTAGGCCAAGGCTGAGTTGTGACCGCAGGGAGAGAGAGAAGGAGCA
CCCACAGCAGGCGAGGGGCTGCGGGAGGCTGTGCTGGGTGGCCGGTGGT
GGGTCTGGGGGCCAGGACCGTGGGAGGCTCGAGGGGGAGCAGGCACGG
GAGGGGGCCCCTGGAGCGGAGAGTCCCTGCTCCAGCTGCCGCCCGACCCC
AGGTCCACCTTCATTTCACAGCCTGGCCCCGGCCGCTCTGACCGGCCCT
GCCCCAGCAGGTGTAGCGGGCAGTGAAGGCCAGGCTCCGGCCGTCCCAA
Contig 74 (450 bp)

GCAGGCTGGCAGCAGGAAATGATCCAGAAAGTGCCACCTCAGCCCCCA
GCCATCTGCCACCCACCTGGAGGCCCTCAUGGGCCGGGCGCCGGGGGCA
GGCGCTATAAAGCCGGCCGGGCCCCAGCCGCCCCAGCCCTCTGGGACCA
CTGCGTTCAGGCGCGCGGCAAGCAGGTCTGTCCCTTGGGCTCCCGTC
AGCTCGGTCTGGGCTGTCTGTGGGGCCAGGCGATCTCGGCAGGAGGAC
GTGGGCTCTCTCTCGGAGCCCTTGGGGGTGAGGCTGGTGGGGGCTGCA
GGTCCCCCTGGGCTGGGCTCAACGCCCGCCGGTCCCGCAGGTCTCACCC
CCGCCATGGGCCCTGTGGAGCGGCTCTGCCCCAGGCTGGGCCCTTGC
TGGCCCCCTTGGAGCACCCCGCCCCGGGCCAAAGCCTTTCATGAACA
Contig 75 (1363 bp)

CCTCCAGCTGGGCCCCGAGGGCACCGTGCCTCTCAGGGGACACCACGGG
GGGCCACAGTGGCTCTCTCTGCTCCAGGCTCTGCTCCCGCTGGGGCCCC
CTGGGCCGCCCGCCCATGGCCAGGGCAAACTCCAGTGCGGCTGCCCGTC
TGGGCAAGAGGGCCGCCAGGCCCGCGTGGTCTTAGCAGGCACTGGCGGA
TGCCGNTAACTAACCATTTCTTCCGAGGAGTCCGAATCTGCTCTGACCA
CGGGCCCCATAAAATCGCTCTTGGCCCCGAGAGGATCCCGAACAGCGGGG
CTGCCCTCTGCTCTCTGCCCCGGCCGCACTCGGCAGGCACGTGCCCTC
GTGCCTCCCGAGTCTGTCAACCGTCCCGTCTTACGATCCCCAGAGTCCCA
CGCGCGGGCAGCTCTTCCACACCCCGACGGCCCCGGAGCTGCCGTGGGC
ACCCAGATCGCCCCCTGACGCCCTTGTCTCTAATTCTGCTGAAATACACAT
AACGTCTCTTGAACGTTTGTCCATTTACGGGGACAATTCTGTGGCCG
TAGGTACACTCCCCCTTGGGGCGCAGCCATCGCACCATCCGCTTCCAGGAG
GTCCCGTCTGCCAGATGGACACTGTCCCCACTGATCCCTAATTCCTGT
CCCCCGAGCCCTGCCCTTCTGTCTCTGTGGCCCTGGCGCCTCCAGGGA
GCCCCCTGTGCGTGGGATCACAAACGTGTGTCCCTTGGCGTCCGGTGTGT
GTCTCTGAGCATCCGAGCTTGGGGTGTCTCCACGCTGCGCCTGTGTGAG
GACGTCTTCCCTTTTGGGCTGCGCGATGCTCCCGTGGGGCTGCCCA
CACTSCGCGTGTTCGCTCATCCATCCACTAAGGCTGAGTTACTTTTGGCG
GTTGTGAATACTGCTGTGTGAACAGGGCGTGCAAAATACCTGTGGAGGC
CATGCTCTTAGGCCTCTCGGGGGGCACACCCAGAGCGGATATGCTCAATA
AGGTAATTCTGTGTTTAGCTTTTGGGGAACCATCAGGCTGGTCTCCAGA
GTGACGGAGCATGCGTCGATTACAGGAATGGTGTCTGAGGCTTTGAGG
TCTCCCACTCGCTTCTATTTTGTGCGTACAGCCGTCGGAACGGC
TGGGTGGTGCCTCTGTGTGGCTTCAATGTGCTTTTCTTTTCTGGCTAT
GAGGTTGAGCGTTTTTATGTAATGTGCTGGCCATTGCGAGGGTTTTGGG
GTTTCTTTTCTTTTGGCTTTGGGGACGGCGCCAGAGCGTATAGAAGT

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FIGURE 6, CONTD.

CGTCGGATTCTGTTAACCAGTGGCCACGACGGGGACCCCCAGGGCTGGC
GTTTCCCTCTGTGTGCACACAGTGGACCTGAGCCAACCAGCAGGGCCTTC
ACCACCACGGCGCAAGAGTCGGCAGCAAGAGAGCAGTGTCTCATGGCTCA
CTTTCTCCCCCTTCCCCGAGTGGTGACAAAACCCCGCGCCACCGGACT
CGGTTAGACAAGGGCGTGGCCAGTGGCCCCGTCTGTACCCCGCACGGCAC
GGCGCTCTCCTTTCTTTCTCGGGGTCCACCACGTCTCCTCAGTTTCCGC
ATGAGAGTACCGCGCTGGCGGGGTGGTGGCTCTGGGGTCCGGGGCGGTG
AGGGCAGGGCTGGGCTGGGGGAGGCAGGTCTTGGCCATTACGCGGGGG
CAGACTCCACATCACACGCTCTCTGTGCTCTTGGCTGCCTGACACATG
GACTTCAACAGGAACAGCGGTGGAGGCATTGACGCCAGGGCCCCGGTT
Contig 82 (550 bp)
TGACACCTCCAGGCAGGAGGTCCAGGCTGGGGTCCAGCTAATGGTGTG
CTGGCTGTGGGGCGTGGGCTCAGCTCTTAGGATGGTGGGCTGGGGCGCG
ACCCAGCAAGGACAGGGTGATGGCAGGTCTGGGCTCAGCAATGAGTGC
CCAGTTTGGGGGTGGGCACTTGGGCTCAGGGGAAGCTCATCAGCTTG
GAGAGGACCGGGGAGGGAGGGGCTTGGCCAGCTGGCCAGATGCCTG
GATGTGAGCACTCAGTGGCCCCGGGTCCACCTCCCTCCAGTGCCATCT
GGGCAGGAGGCTCCGATCCCTGTCCCTGGGACCCGCTGTCTGAAATGAG
GTTCACTTGGTGCCTTCCCCAGAGATGCTCGGTCCGGAAGCTCAGGAGGC
AGGAGTGACAAAGGTCTGGGGAATGGAGCAGAGTGGGCTGGGGCACA
GAGGCTGCCCCAGCCTGGGAAGATGGGGAGCTTTCAGGGGTACCCCGC
CAGCTTCTGGGGCTCTGGATACCCAAGGGTGTGAAGAGGTGAACAGCCA
Contig 83 (984 bp)
CTGAGCCAGCTATCTAGATTAGACCCCGGTCCGTCCCAATTCTTCTCA
AAGCTGTCCCGAGATGAGAGATGAGGTTTTCGTGTCTCTGTCTCTCTCG
CTTCCCTGGGATGTGCCCTAGGTTGGGAGAGGGTGTGTCCAGGGCTCA
GCAGGCGGTCCCATCTTCCCGAGACGGGAGAGATCCCTCTCTCGGG
CCTGTCCCCACGGCCCCACACACACCCCCCCCCCGGATGGCACCCAT
GCACCTGCCATCTGTCCAGTAGGGGATGGGTTTGGCGAGACTGGAGATG
GCTGTAGCCACTGAGACATGCCCTGCCACGTAGCTGACCCCTGGGTGT
GCTGTGTGAGATCTGGGGACCCCGACACCTAGGGATCATCTTGTGCCA
GCTTCTTGGGACCTCTCAGAAATGGGGGCCCCAGAGGGTGGCAAG
GTGATGGGAGGCTGGGAAGTCTGGCGGTGGCGGGGTGGGTGGGGGCA
GTGCGGGTGGGTGGGGGTGTCTCGGGGTGGGAAGTGTCTCAGCAAGGT
TTTGGACACAAAGTCAGGAGGAAGGATGACGAGGAGACTTGCAGAATTA
CAGGTAGAAATCAGGAACCCACATCGACGCCAATGATCTATCCCCCCTT
TGATTGTTTTCTCTGGGGCTTTTTTCNTTTTTTTTTTTTTTTTTTT
TTAATCCCTCTTAGCTTTTACGCGCTCAACACCAAAATTAACGTACTC
CCCACCCACGTAACAGGGGGGGGTGACCCGAAGGACGAGGAGCACAG
AAGCCACCATCCCTCAGCTTGGCGGCACAGCCGCTGTCTGCCCTTCCCC
CCATTATCGCCCTTGAATTGATTTTGTCTTGTCTGTCTCTGTCTGTCT
GGGTAGAGTGGAAAAGGGAACTCTGTGGGGGTGCCAGCCACTGGGCCCC
CCAAAGATTTCAGGGGAATGAACGGGTGCCGCC
Contig 84 (550 bp)
TGCCCTCACAACCTGCCCTGTAGCCACACTCGCGACTAATAAGGCGA
GAGGTCAGCGGGCAGCCCCACGGGGAGAAAGTGCCTCCGTGCCCCACC
CCTGGCTCTGATGGCCAGCCTGGCAGCCCAAGGTGGCTCGGCTTCTCT
ACCTCCAAGGTCCAGGCGCATGTCCAGCACCAGCAGAAGCTTCTCCAG
GTTGGTGCCTGTCTCAGGGCAGAAAGCAGGGGTGAGGCTCCCCAAAGGGCC
ACTGGCACCAATGCCCCAGGCAGCCAGCGAAGGGGACAGCCACCCC
CAGCCCGGGGACGAGGCTGAGGGGACATGGGGAAACCCAGAGCAGGGCC
AAGGGGAGCAGAGCCCTCTCTCGGGACTTGAATCTTTCCGGGGGGCC
CAGGGAGCTGGGGTCTGAGAGGGCTTTCAAAATACGGCCACCCCA
AATTGCCACGTGGGCCACAGAGCAAGGAGTCCGTGCCAAAGTGGCTGGC
TTCAGCGCAGGAAGTCCCTCTGGGGCTCCCTCTATAGGCACAGG
Contig 85 (500 bp)
TGAGCCAGGGCTGGCCAGCTAAGCCCTGGAGCCCTCCCGCCCTGTTT
CCTGCCTCCATGCTGGCGGAGCTCGGCTTACTGAGCGGGGCCAGGCCA
GTGTGCGTGTGGAGGTAGATTCCACTCAGCTGGAGGTTGAGTGGGCAGG
GGGCGCAGACCTCAGGCCAGCTCTGSCCGGCCAGGTCTCTGAAGCTCC
CCCGGCTGGCTCCCGCTCCCTGCTCTGGCTTGTCTGGCCCTTGGCT
GACAAAGCTTCTGTGCTCTGCTCTCAGGAGAGACACTGGCTCCCCCGCT
TCGGATGAGGACGGGGCTTTCTSCACAAGTCTGCCCCAGAATGTTGG
GGCGCCAGCAGCTGAGCCAGCAGCTCTCCCCCTGCCCTGGCTGGACAC

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FIGURE 6, CONTD.

GAATCCCGGCATCGAGGCGGGAAGGGGGATGGAGGGATGGGGCTACCCA
CCCCTGCTCCCCACCCAGAATAGCTGGGCGGCCCCATGGGAGGCGCGCC
Contig 86 (913 bp)
CTGTTTTCACGTCTTCTGAGGACACACCCAGAAGAGGGGCTGCAGGCGCC
CATGGTGACTCCATGTGTTCACTGCTGAGGCCTCTGCAGACCGTCTCCCCG
CAGCAGCCGACCCGTTTCCATGCCACCAACAGCGTGCGAGGCGCGCACTG
TCCCCACGGCTGTGCAACTGTTTGAATCTGAGTTATATAAGCAACAGAC
GCTCCTTCAAACACACTCACGTGCACAGTGGCGCACAGGCGCACAGACAC
ACACACGGAGTAATAGGCCTCCCCCCTCCCTGAGCCAGAGGGGGGCT
GGGGCCCTGGAGCCTGTGCTTTAGGGCCTTTAGGAAGCTGGTGCTCC
CAGAGGGGCGCGCCGAGCGTTGGCTTCCCAAGTCCCCACCAACCTCGA
CAGACTCAAACGTTGGTTTCTTCTGCTTTTGCCCAAGGGATGGGCGCCG
AGGTGGCCCTGCTGAGGTTTACGCCAGCGGCCAGGCAACCTTCTCT
CCCGTCCCGGCCACTTCAATGSGACAGCGGGCTTCCCCACGTTGTCC
CCTGGGTTGTGCTGCTTTTCGTAATGAGACGGAGGCGAGCTCCACCTGTCC
TGGGGTGAATCTCTTCTGCAAGAACTCGCTTCCCGCGGCTGGTCTGT
CTCTTCTCGGTGTGTGAACCTCTCGTCACCAAGAAAGGTGGCTGTGAC
GTGCGCCTTCCCTCGTGGCTTTTGAGTCTGGGTCTTGTGCGGAACC
TGCCCCAAGAGGGGAGTGACCCCCACGAGGGAGAGCTAGCTCCTGTGG
CGAGCAGACCGGGGCGCCCGAGATTATGGGGTTCAGCTCACAGTCGCA
TSACGCTGCCTTTGGACGAGGCGAGTCAAGGGAAGCTTTTCTGCCA
CGAGCCACAGGCA

Contig 87 (650 bp)
TCCACACCTGTGGAGCGCTGCCTCGCTGATGCCCTCTGCCAGCTGATG
CTCAGGTGCCAGACTTGGGCTCAGTCCAAACAGGGGCCACAGGTGCT
GCACCTGGGCAAGGGAGCCTGTGCGCAGGGCCTCAGGTCTCCAGGCTCG
CTGGGACCGAAGCGCACTGGGTCTGGACTCCGGGCTTCCCCAGGGGCTG
CTCGGGGACACCTGGAAATGAAGCCCCCTGGCTCATAGGTTCCACGTG
AGGGCCCTGAGGCCACCAAGCCACCAAACTCAGTTAAGCGAGGGAG
CTTGGGGCTGCTAAGCTCCAAGCGGAAGCGGCGCACTACGACTGCCCT
CTCTGCCAGCCAGCGCCAGCTTGTGACGTCCCAACAGGCCAGGGAC
CCTGTCCACAGATGCTGGGCCCTTCACTCTCTGCTCCCTGGAAGCGCT
GGGCACTGTGTGGGCACACAGCCCGCACCCGCTGTAAAGGAAGGAAAGG
CCCCATCTCAAAAAGCCGTGGGCGAGTGGGCCATGATGGTCTCCGAG
GCAGGTCTCTTGGGACCCCTTCTCTCTCGGCTGCCCCAGGAGCGGCC
AGGTCTGCCCTGGATTAACTCTGCCCGCATGTATTTCAAACTGGCTT
Contig 88 (700 bp)
TGGGGCCCTTTGGGGCCGGAGCGGCCAGTCTGCTGGGCCGGGAGCAGGG
GGTCTCTGTCCGAGGGAGGGGGCTGGTCTCAGGGAGGAGAGGAGGCA
GGTCTCACCTGAAAGGATCTGCCCTTCTCCTCAGGCTCTGGGATGCTGG
GCAGAGAAACAGAGGAAAGGCCCAACTTGTGGCTGGTGGGATGGGG
CCGGGGTCTGCTCCCGGCACACCCCGCCCAACCCACCTTAGTGGCCAA
AGTGGGTGTCATGATGGCCACTGACCTCACGGGGCGCAGGAGACAA
AATTTCAGCCACTCTTGGGGGAGGACACTTGTGGCTGAGTCTTAGGGG
CTGAGTTTCCGGGGGGACCCAGCTCTCCCCCAGTATCAGACACCCCTG
CCCACTCTCCACGCTGCTCCCCAACCCAGTGTCTTGGACGGGCATCT
CCCCGTGCCCTGCAGCCGCTGCTCTGACCATGTCCCTCCCCACCT
CCCCCTCTGCAGGGCCAGGCTCCAGGAGCAGAGCCGAGGCCCCACCCCTA
GACTGAGCTGGGGACCGAGACCCCAAGTCGCCACCCGGTCTCTGCGTTAG
AGAGGGGTTCGGGGGGCACCCCTGGGGCGGCAC'TGGGGGGCGGGAAGGA
GAGCCCTGGGCGGT'TCTGGGAAGGCTTGGGAGGGAGGGAGGGT'TTTC
Contig 89 (1400 bp)
GCACACCCGGAGAACAGAGGGAGGGGCTTACCAGTCTCAGGGTTTTTT
TGGGGATTCTTTGAACCTTGCCCTATTGGTTTCGAGGCTTCTGTTCTCTC
CAATCCCCCTTCTGAACCCUCCAAAAATGGGTTACGCCCCACCCAG
CCAGAGGAAACCAATTGGGGGATTGGGGGGAGGCGGGCCAGCAAAAGCC
TTGGGCCCCAGCCCCCTGGCTTTTGGCTCTGGCTGCCAGGTAGGGGG
AGGGACCGGGTGACCTCCGGGGGCTGGCCACGGACTCTGCCUCCACCCC
CAGGGCAGACGTGCACAGGAGCGGAGAGGCTCCGAGGAATGAGGCCATCA
AAGGGACAGGTGAGGCCACGAGCGCTGGGACCTGGAAGTGT'TAGGGCCT
GGGGGACGAGGCTGCGGCTGCGGGCTCCGTGGTCAGGAGGCCCTCTGCC
CACTGAGCAGTCCCACCACTGGCACACGAGCCTCTCTGGGGTCCGGCTG

FIGURE 6, CONTD.

CTCTCCGGCAGGGGTGGGCTCTGAACGTCCAGCTCCGCAGACAAATCAGA
TTCCTCCGAGCCCTGAGAAAGCCCTCCCTCCAGCCCGTCTCCACCTG
TCGGTGGACAGAGTGACCCCTGCTGACCCCTGCCCGGGCTCCCGCAGGA
GATGTGAGAGAGTAAGAGGCGGTACAGGACGGCCGGGGCGGCCCGGCGA
GGTGCAGGTGTGTGGGTGTGAGGCTGGGCACAGGCTGGCACAGCCTCCCT
GGCCAGTCCCTTGGGCACCTCTGGGCACCTCGGTGTGCTGCCTCCTGA
AGGGATCCACCTCCAGCCACCTCCTCTCGGGCCAGCCCCACCCACCC
CCGAGCTACAGATGCCCTGCCATTCCGCCCCAAGTGTCTGGACCTGGAG
CCAGGCAGCCACCCGCTCAGCCTGGCCAGACCCAGCGTTGCCCTTCAGC
CCCTCCTCCTCCCGCGGGTCTCCGCTCGTCTCCTCAGGTGGAAAGC
CCCTTCCACCTGCCATCTTGCTGCGCCAGGATACACCGCTCAACTCA
AGGCTCCTCTCGCCCTCTCAAGGCTCTGTCCAGGCCCTCTCTGAC
CTGGCACCACCTGCCCTCCTGGCAGCCCGAGCAACCCCTGCCACAG
TCCACCACAGTCTCTTCTGGTCTGCCCGCAGGATGCTTCTAGAACTGG
GGGGGGGGTCTTCCAGCCACGCAGCATCCACTGGGCCCTGGGGTCCCT
CCCGAGGTGCCCTCAGAGCTTGACAGTGGTGCAGACGGCTCTGTCCGA
ACCATGCTCCCTGCCCTTGAGCTCGTGAGATGTTCAGGTCATTTG
GCTGCACCCAAAGAGTGGCCCTCAGGGTCCCTCGGCCCTCCATC

Contig 90 (350 bp)

GTACTGTAGGGCTCATTGGAATAGCTACTAGGTACAGCTGATCCACA
CCTTAGGCCATCACAACTTCCAGAGGTAGTGCCGCTCCTGTCTGTGAAC
AAGACGGTAGTGAAGTGTGTGAGAGCTCAGATCTGTGGGTCACTGACCG
AGTGTGGAACCTGGGGGAAGGCTGTGGGTGTCCCGGCTGGGTGGCCA
TGTATGTGCCCTTTCTATCCCTTGGACGAGGCTGGTTCACTCGGCTCT
AGAGCCCCAAGCCCGAGTGTCTGCCAACCCCAAGGCTGAGCCTCAT
CAGACCCACCACTCATCGCCATGGCTACGCAGGACACACCGCTCTCCAC
CCCCACAGCCGCCACCTCCCGAGGTTCCAAAGCTTGA

Contig 91 (1464 bp)

TCCAGGACCTGATGCAGCAGCCACGTCCGAGAGCCCTCCACAGAGGCC
CTTGTGACCAAGGCTAGGGGAAGGGCACCAGGGAGATGCTGAGAAGGGG
CCTTCCGAGGGGGCAGGTGGGACTGACTGTGACCCAACTCCCAACCC
CCTCTCCCGCTCCAGAGGGTGCCAGCCTGGAAGCTGCAAACTCCAAATCC
ACAGGTGGGCTCAGTGGGGAGGCTGGTGGCCCCACCTGTGGGGCCCC
AGCTGCTCTGGGCGGGGTGGGGCTGCTCCAGCAGGGTCCCATCCAG
CTTCTCCCTGGGGAGACTCACAGTTCTGGGAGAAGGCTCCTGACTGCACC
GCAGCGCCCCCCTCCACAGACTCACCAAGTTCTCTCTGTCATCGG
TGAATGCTCTCCGCAATTTGCCAGGCTGGCATCTCCAGAGGATACGT
CCAAAGCCAGGGCAAAGCCGGGCCCTGCCCGGAGCTCCCAACAGGCG
TGAGGGCTGGGCTGGATCTCGGGGGGTGGAGGGGAGACTCAGAAGGTG
CAGCGGGGTGGAGCGAGGCTGAGCCAAAGTGCACCGAGGGCCAGAGAG
GCCGAGGGCGGCAGGAGGAGAGGCCAGCCTGGAGGGGGTGGGTGCC
CTGGGCAGGTCTGGGGCTCAAGAAGAAGAGAGTGTGTGTCAGGGGGCTG
TCCAAGCTGCCCGGGAGGCTGCTGCCACCTCCAGGGAGCAAGCAGGG
AGGCTGCAGCTGCCCGGCCCGCCGCTCTCCAGGACCACGGCTGGCCCCAG
GCCTCAACGCTCTCCACAGCCAGGAGACCCAGGGCACCCGCTCCATT
TACCGCGGCTCCGGGTCCGTTTGCTGCGCCCTGGGATGGACTGTGGGG
GCGGGGCGCTGTCTGGGGAGGAGGAGGTGTCTGAGGCTGGACACCTTGA
AGGCAGGTGAGAGTGACAGGTCCGTGCGCAGCAGCCTTCGGCTCTGGATT
CTGGCCCTGAGCGAGGGGCTGGCTGGAAGTGGGCCGGGGCTGCCCGAGG
AGAGTGTGAGGGAGAGGAGACGGGTTTGGCCCCGAGGTGCCGGCGTG
GTGCCCTGGAGTGGGCTGAGCGGGAAGTGGGTGTTGGGCTCTGGAGACG
GGGGTCTGTGGCTTGGGATGGTGACAAGACCCCTCAGGTGGAGGCGGCC
GCAGAGGAGGACAGAAAGCCAGGCCCGCCAGCCCGGGGAGGCTGGG
AGTCAGGAGGGACAGCAGAGCCCTGGGCTCAGTGTACCCGCTCTGGCA
CCTCGCCGACGGATGTCTGGCCCTGCAGTGGTTGTCCCTCACCTGAG
CCCTGAGAACCATGCAGGATGCTGGTGTACAGCAGGAGAGCGCCAGGGC
CTGGGAGGAGTCTTACTGGAAGCCCTTCTCTTCCGTTTGACAGCAGGCG
GGAATGACTGGGGG

Contig 92 (694 bp)

TGGAGCCAGGGCACGCGCAGAGCGGTCCGAGGCCGTGCGTGTGACCCGG
GGGATGGGCGGACCTGGGGGTGGGCTGTGAGCCAGGCATAGGGACCCCG

SUBSTITUTE SHEET (RULE 26)

FIGURE 6, CONTD.

ACTTGGGCACGGCCAGGTGGGGCCGGGCAAGGGGGAACAAGGACGCTGGC
CTCCAAGGGCCCCACGTGGGCACAGAGGAAGAGCCGACCCAGGTTGTGGG
CGCATGGAAACCCCCCACTCTGGGGCCAGGAGGCCGAACGTCUCAAAGGGC
TGAGGCTGGGAGGGAAGAGTCCCTTTGGGGGTGAGTCAGTGTCCCTTGTG
GGTCCCCCTGCCACTGGCGGCACCTCTGACCCCACTCCTTGGGGGTG
GACGGTGGATGGATTCTCTGCAGCCTTCTTCTGGAATAGTCTCTGCCAT
CCTCGGCAAGCAGTGATTGCTCTGCCAAGTCCAGGCCCGCCCTGCAA
GGTGCTCCCAACCAATGAGCCCCGGACAGTTCGAGGGCTTCTCAGCG
TACTGAGGGGTATGAACAGCTGTCCCTCGGAAAGTGGGGACAGGCC
CTGCCACTCCATCTCGGACGCCCGGTCTAGTCAGCACTTGTCTCCCTG
CCTTGTGCCCTGACCTTTTGTAGGACCATCAAACCTCAGCCTCTG
CCCCAGGAGGTCAAGCCCCCGTCCCGACGCCUCCAGACCAGCA
Contig 93 (900 bp)
CCAGCCCCATCCCCGGCTGGTCCCCACACACAGAGCCCCCGTTTCC
AGGGGACAGCACAGCCTGCCCCAGGTCTTACATAAAGTCACCTTCTCAG
AGCTCTCTGTGCGGGCTCAGGGGAATGAATCTGACAGCATCCATGAGGAC
ACAGGTTTGATCCAGGCCCGCTCAGCAGGTTAAGGATCTGGCGTTGCC
GTGAGCTGTGTGGAGGTGCAAGACGTGGCTCAGATCTGGTGGGCTGT
GACTGAGGTGGCGGCAGCAGCTGCAGCTCTGATTGGACCCCTAGCCTGG
GAACCTCCATATGCCGCGGGTGACCCCTGAAGGACAAAAATAAATAA
TAATAAAGAGTAACACACCTTCTTAGCCATAACCACTTCCCTAGG
GGCGGAGGGCCAGGAAGCGGCACCCCGCCCGAGGCTGCCCGTGCGCC
CGGGCAGGCGGCTCAGCCTGCTTTTGTCTGTGATGTGAGCCGCCACGC
CCCACATGAGGGGCTGGGCTGGCGAGTAAGTCTTAACTGACGGGAGC
TTGACACAGCAATTCACAGCGGGGATGCAGCCGGGAAGGAAGTTATTC
GTGTGTAGCTATTAGGCGCGGAGTGAGGGTGTCCCTGCCCTGGGCCCA
CCCTTGGGGGAGGCATCACAGGGGTTTGAACACCTGCCATGAACACG
GGGCAAAAGCCAGCCAGGGGCGAGGTGCCTGAGGCTGGGAACCAACCG
TGTCTCTGAAATCCGGGGAATGCCCACTGCAGGCATGTTCAAGCGTCAA
GACCGGGGCTCTCTGTGAGAAGGACTGGCGAAGGCCAATCAAAAAGCGC
ACCCCTCTGTGCAAAACCCCAACCAATGGAACAAATCCAGAGGGGCCA
Contig 94 (550 bp)
AGTCTCGGCTGTGTCCATGGGGTTGCCAAGGTGCCAGGCAGAGACCTTGG
GGACAAAGCTCTCTGAGCAGAGGACATGGCCACGTCCCTGCTCAGCA
GGTGCACAGGCTGSEGTCTGATGCCCTCGCTGGGGTGGGGCGGGTTGAG
GGGCGAGGCCAGACACCTTCTGTCCTGCGGAGTTGTTTCCCTTCTG
TTCCTGGAAGCCCCCTGACGTACAGGAGGCCCTGGGGCTGACGCTG
CACCTTCTGACACCTGTCTCTTGGGGATGGGACAGGACAGGGAGACCCC
GGGGCTGGAGCGGAGCGGTAGACAGAGAGTTGACTCTCTCTGAGTCT
GTGAGGGCTCTTCCCGGCTTGGCTTCTGTCAGGGGCTTTCGGGTCA
GGGTGGCTCAAGGTGACGAAGACCTGGTCTCGGGAGTCTGACAGCGCA
AAGTTGACAGCCACCCCGGGGAGGGCGGCCAAGGACAGGAGGGCC
CAGGGAAGTCTGGGCTTGCAGGCGCTCCGGGTGGGGAGGCCAAGGT
Contig 95 (1200 bp)
GTTTGTCTCAGCAGGCAAGGCTCCGAGGCCTTAATAGCCCATATGA
CAGCGCCGCTCTTGGCATGGGGCCCCCTGGCATGGGGCAGGGCAGGG
CAGAGCAAGCAGCATGCAGCTTCTACCTTCTTCTGACCTCTGTGGCCCT
TCCGAGGCTCAGGGGCTCCCCGAGTGGGACCCAGCCCTGGCTCTCT
TCCAGAGCCAGGCCAAGGCTGGGAGTGGCCAGAGATGAGGGTGCCG
AGCAGGGCACTGCCTTGGCGTCCCCATCCCTGGCGCTCAGGGCGTACT
GTCCAAAACCAAGAAACAGTCAGCAAACTTCTCCAGCAAGCTGGG
GTCAAAGGTGGCTTCCGAGGCGTGATCAGGTTGGCTTTGCTACTGTAC
CGTGTGCCCTGGCAAGGCACAGGGACACAGACACACCTCCGAGAACC
TGGGGCTTCCAGGGGCTCAGGCTGCTGGGCCATCCCGGGCCCCCTGTGGT
CCCAGGATCTGCCGGGACCGTGAGGCTGCTGCCACCTCTGCTTGGGA
CAGGCCCCACAGAGTTCACAGCCAGGGGACCGGGACAGGGCCCCGCTG
GGCACCTGCTCCAGCCTCACCAAGCTGGGCCCCAGGCTGTGCTTGC
GACACCTGAGTCTCAGGACGGGCGGGGACAAAGCCGCCCGCCCTCC
CCCCGCTGGGAGGAGACCCGCTGGCCCTGACGTGTGGGCTGTGAGAGC
TGAAATETCACAGCAATTAGCCCTAACGAGGCCGAGGGAGGGAGCGGG
GGAGGCGGGGAGGGGATCCACGAGCCGAGGGGCCGAGCTGCCACCC
CAGCGGTGATTCAGGCACTCAGGATTAATTCGGTCTTTAGAACTCAGC
CGGACGAGAGAGCGGGCCAGGCGGGCTGTGCCCCCTCCACCGCCCC
TTAACAGGTGCCCGAACACGAGGCTGTGGGAGATGCTGAGGTGCGCAAG

SUBSTITUTE SHEET (RULE 26)

FIGURE 6, CONTD.

AAGAAGATGCAGGAAATCTCAAAGTTCAGTCACAAGAAAACCAATTCA
AAAACCAGCAGAGCAGACATACGATGGCAAATAACCACGAGAAAGTCAGC
ACCCGCTGTCCCTGGGGGGACGCGAGTCAAAGCCAGGAGACACCAGGAT
ATGCCACTGCCAAGGCTACGGATAACGGGAAGCAAGAGACACAGACAGA
AAGGATGCTTCGGTGTGGGGAGGGTGGGGTGGGGCGGGGGTCCCCC
TGGAGCAGGATGTGAAGGCACTTGGGGGGGCTTGCACTCTGGGGGCC
TTTGGCACAGGCGGAGGGCCCGGAAGGCTCTAGGGGCACGAGAGGGGT
GCCAGGCTTCTTACCCAGCCAGGCAGACCCAGCCCTGTATGAAGCTT
GACGTGCAGCAGCAAGAGCAACATGCTACAGACATGTGTCTGTGTGTG
TGTG

Contig 99 (1000 bp)

CGTTCTCAGGCGCACGGGGCAGAGGCTGACGGTCCGAGGGGCTTTGGGTG
CTGGAAAGCCTGAGTTTGAATCCCAGCTCGGTTTCTTAAAGCTGTGTCTC
CAGCGCCAAAGGAATGGGGCTTCTGGGAAAGTCTCGGCTGAGGCTGGC
GGGACCTGCCAGCCCCGGAGGGCATCTGACCAGACAGCTTCTCAAGCTCA
CAGGGCTTCAATGGCAGGATGGGGAAGGCTGTGGTGGGAGTGGGGAGCAC
TGCACACCCCTGCCAGGCTTCTAGTACGGTGGCTCTCAAAAGGGGT
TCTCTGTGTCCAATGAGCAAGTCTTTGTCCGGGGCAGGATTACTAAGTCC
AAGGGTGTCTGCCCTCCCTGGGGCACAGAGCAGGGGCCAGATCACGT
GGCTGTAACCTGCCAGGTGCAAGCTGCCACCATGTCCCACTGGGTCT
CCAGTTACCTTGGGAGGTGCAAGGTGGGTGATGGGAACTGAGGCAGA
GAGCTGGCAAAAGAGTGCCGGCAGGACTGCCGCCAGACCCAGCTAA
CCGACCCCTCACACGGAGTGTCTACTTTGCAGCTTGGACGTGGGAAAA
GGTTACCCACAGCAGCGTGTGCAGGCACGCTGATGTCTGTCTACTTA
TGCATATGTTCTACGTGCATGCACGTGAGTGTGTCTGTGTGCTTGTGCT
GT
TGAATGCTTGT
GATGT
TTTATACCTGT
ACGTGAGAAATGTGCACTCGTGCATGTTTGCATGTGAGTTTCAATGTACAC
TGCTTTTAAAGTGTGCAGCTGTGCACATGTGTCTGTGTCTCTTGCACG

Contig 100 (1500 bp)

CGTATAAATATATTAATATAGAAATAAATAGATTGATAATATAGATAAAC
TAAACCATTATCAATACCGGGTGGCCCCAGCAAGGATACTAGCCAGTT
TATCAAGGTGCTAAGTCAGCAGATAGAAATGGCCACAAACGAAACCTGTA
CTGCCATGTCCACTCTAATGCAATATGCCACTGACATCAGTGGTAGGTG
AGCTGAGTCCATCTGGGCTCCCACTTCCGGGCCCGGCTTGTCCCAACGG
AGGTTCCTTCCAGGTTTCCCAACCAACCGGGCCCCAGGTCTCCCTG
TCTTGACTCGTTTCTGGAGTCTTCTGGGGCTCTGCACTCTCCCTTGTG
GGGCTTCTGTCCCTTCCCTTGGGCTTGGGGCTTGGGCTTGGGCTTGGG
TCCCGGGCTTGGGGCTTCCCTCTCTTCTTCCCTGGAAGAGAGGGAGCC
AGGCTGGGCGGGCCAGGAGGGAATGCCCTGACTCTGCTCCAGATGGAC
AGGTCCGGACATGCACTGGGCTTCCCTTGGGCTGCTGAGCCAGAGCAGG
ACGGGTCTTCTTGGAAATCTGGGGCCAGCCAGGTTACGCTGTGGGTGGG
CAGCCGCCAGCATCTGTAGGGGCGCTGACGGCGGGGAATGACCTCGA
CTTCTGCTTGGCACCCAGCTCTGGAACAGCCCCCTGGCGAGCTTCCGCC
AGAGCTGGGCCAGAGGTTCCCTGTGCGGGGACCCAGCAGGGCCCTC
CCTGACTCTCAACCCACCTGCTGAGGAGTGGCCCCCTGGCTCCGT
GGATCTCTGGGTGGGGCTCAGCGGCTTGCAGGCTGGGAACAGCCAA
GCACATCCCCAGGCTTGGCCACACCTTCCACCGGAGCGGGGATCTG
CATTTCCGAGGCTCTGGGGCAGCTCTGAGAGCCCCGGGTCTCGAGCC
CAGCCGTGGCGTTGTACGCCCTGGGGGCTGTGGACAGCGTGTCTCATT
GCCCCCTCGAGGTCCGGCCAGGTTCCCTCCACCTGCTCGCCAGAGCC
CTCTCCCAACCAACCACTTCTGTGTCTTGAAGCGGGACACCACT
CCGGTTTACGGACCTTGTACGTGCGGCTTCTCTGAGAGAAATGCCTG
GAGCAGATGTTTGTCCGCAGGCTGCTCCGGAGGCTTACCAGAGCCCC
TCACCTAAACGGCCGGGCTCAGCAGCCGGGGCCCTGTCCCAACCGCCC
AGGTGGTGGGTTCTCTGTGCGAGTGTGGGCATCTGTGAAGATACTGT
TTATCTGCTCATCGTCTGGTCTCCCCAGAGGTAGAGCAGGGCCCGGCA
CAGCCGTCTCGGGTGGGCACTGCCCTTGGGGCTCAGGCTCCATGCA
GGAGGACGCTGTTGACAGGAGGCCCCGTGTGAGTGTGGGGGCGGCC
AGCCTGCCTTAGGTACAGCCAAAGCGGCATTAACCACAGGCCCCCGA

FIGURE 6, CONTD.

Contig 101 (600 bp)

TCTAGAATACCTGGCCCTCCAGGGACGTGTCTGTAGCTGCGGCTTTCAG
GGCAAAGTGTAAATTAACATCCCCAGGCTTCCCTTCCAGTTGGCACAGGG
CACCACATGAGGAGCAGCCTCTGGGTGCCAAAGGGCCCACTGGTGGCAG
GCGCTGGGCTGAGTGCACCCCGCATGCTTCCCGCCCACTCACCTGCTGG
CCCCACCCCTGACCACAGCACCTGTGGGAACACTAGGCTGGCAGCCACA
CGCTGCTCTCACTGGAGGCCAGTGCCAGGCAGCCTGCTTGGCTAGCCTAG
CAGATGCCCGCTCGCCTCTGCCCTTGGCCCTAGCCCATGCAGGAGCCAG
GTTGGGGCACAGGAAGGACGATTGGGGCCCCAGGTACGGCACATCCAGGC
CACAGCCGTGGCCACACGAAGGCGGCCCTGAGGGGGCGTTGGGGGGCAGA
CCCTGCCCCCGCTGGCCGCCAGCTCCAGGCATTAAATTCACAGGGACC
TGTTCAGCTGGGTGGCCGCCAGCCTGCCCTTGGCTTCCAAGGCCCTTA
AAATGCCCTCTTTTCGTAAACTAGGACTTACCAAGCTCAGCGAGCCCTT

Contig 102 (1867 bp)

AGTATATCGGGTGAGACTGGGACCGGTCTGCGGGGAAGCCCCACCATAA
AGGCCACGTTGGGCCACAGTCCGGGCCACGTGAGTGTGGCGGGTCCCGG
GGTCTGCTCTTGAACACACAGGATCTCTAAGAGGTACCAAGCCGAGGCCAA
CTTCACGTGAGCAAGTGAACAAATGACTGAATGAGAGCCTGAGCGAATGA
GTGAGGGGTGAGTCCGTCCACACAGCAGCCTAGGCTCAGGCAACCGCTGT
UCCCGCTCTCCACTGGTGACAGAACGAAAGAGTGGGAAAGAGTGGT
TGTCTCCACAACCCAGTCTCCCAACCCCTTGACGCCCAACCCCTCCAG
GGGTGCCGGGCTGGCCTGTGGGCCCAAGTCTGGAGGCTCTGGCACCTTC
CTCATCGCTTCTCCAGCACCCAGGTTCTGTGCTGAGCCCTCTGGCCCA
CAGGCCCTCGGGGACAAAGAGGGCCACCTGGAGGCTCAGGGAGCCTCACCT
GCCTCGTGTCTTGGCGGAGGCGGCTCTGGACATGTGATAGACCGGCTG
GGCTCAGCAGCTCTGCTGGAAGATGTCAGGGACAGCCTGGGCCACTCTC
CCACCAGGAGAACTTATCTCGGTGGGGTCCCCCGGGGAAGGGATGGC
ATCCCCAGCGGGGACCCAGAGGCTCCAGCACACCGAUCCTGTCCCTCCAGC
CCCTGCCCCACACGATGCTCACAGCTCAGCCTCGAACACCGCACCTTGT
GACTTTGCTCTCTGAGGCTGTCTCTCAGCCGACGCGGGCTCCGCTGCA
TGGTCTGGAACCCAGTGGGACTCGGTGGTGACAGGGAACAGGGGCTCTT
GGAGTGGGGTGGCGGGGAGCCCGAGGGAGCTGCTTGGGCTTTGATGG
CTGAGTGGGTGAAGTCAGGCAAGCTCCCCAGGGCTCCCTGACCCCCC
CACCTCAAAAAATCCAGAGCATCCTTTCTTTGGGTCTGGTGAGGCTCTC
TGAGGTGAGACCTGCGTGTCTGGGCAAGTGGGCTGGAGCAGGAAGAAA
GCAGGACACCCUCCGCCCTGGCCAGACTCCCCAACCCAGCAGGAGAC
ACCTGAAACGGGATGGAACCATCTGAAAGAGCCACCTCTCTCTCTTA
TGATCAGCTGCCGGGTCTGGGGGCCCGCCAGGCCCCAGATGTCCGG
GCTGCTCCGCTCTACATCCAGGGTTTCTGGGCTCAGGACTCTGTCCCC
CCAAGCATGCAGGGTCCAGGCTGGGCTCTTCATGCTGCCCGTGTGCA
TGGTGGGAAGGAAGGGACAGTCTGGAGACCCCGCCCTCCCATGCG
TGGCGCCGGGGACAAAGCCGGCTGGGCTCTCAGTTTGGGTTCAGAGCA
AACGTTGATCTGACCTGGTCTGAGATGCTCGGCCGATGCTGCGTTGTC
CGCTCGCATTTTCTCTTTCTCTGGGAGGCGTGGCTGCGCTGTGGCTT
CCGGCCAGCCUACCGAGGGACGAGGCTGGCTGCGGGGTCTGGGGGCC
CCTGCCCGACACAGACGCTCTGGCTCAGCTTTTGTCTCTGTGACCCATC
ACTAAGGGCCACCTCTGACCCGGAGCCCTGTCTCCGAGGTGGGAATTGG
GGGCTGTCCCTGGCTCATAGGACCTGTTGGGGGATCCAGGGCTGTGT
CATGCCCTCCCCAGAACTCTGGGGGTGGCGGAGGGTTTCCCCAGCT
TCGGGCTAGCTGGGGAGGCGGAAGGCGCTGAGGCTTGCCTGTCCCA
GGGAGCATGGCTTCTGCTGCAGACTGGGGCCCCGACACCCAGCCACCT
GGCGCTCTGAAGCACT

Contig 103 (650 bp)

GTTGAGGATTCTCGGCAATTTCTCTGCTACTGGCGCTCCAATCGCCTCG
ATGGGCTTCTCTCCAGATACAGCTGCAGATCTTGGCGGGCACACCGTT
GAGCGTCACTCTGTAGTGCAGATTGCACTCGTTGTCAATGGACATCCAGG
CCATGCCGACGGCATGTGGATTCTGTGCATCCGTGTGCTCTGTGCTTTC
AGCAGAAATGGGTTCCGCCGAGTCCCGAGCATCGGCCACTGGACGGGGCAC
TAGGCGGCCACGGATCAGGCTCGTCTCATGCTCGGTGGCCACATTAAAGC
CCAGTTCCCCGGCATAACAGGACTCGAGGACCTTGGGACCCAATTTCTCC
ACACTACCAATGGCTGGTTGAAGTTGAAGCTCGGCGTCAAGATCTCCAG
CTTGGCTTCCGCTTGGCTGCTCTCAATCAAAGTATGTTGGGCTAT
CCCGGGTGTTCAGTGTCTCCGTTTCGATGTTGTAGGCCAGAGATCCATCG
GTGTTCAAGTAGACCCACGCCAAACCGCTGCTTGGTTCGAGGATTCCGG

FIGURE 6, CONTD.

ACTGTGCGGGCCAGCAGGGTCTGGAAGATTTCGCAGCTGGCTCGGGTCA
CGATGTGTCCTGGATGCGCAGATGTGGGTACTTCTTGGACTCCACGGTC
Contig 104 (1630 bp)
GGTGTGTCACTGCTGTGGCTCAGACCCCTGCTGTGGCACAGGGTCCATC
CTTAGCCCAAGAACTTGCACATGCCACAGGTGCAGCCAAAAGAAAATTCT
TACTAATAAGTTGTTTCAATTTGCCCTTTACGTAGAGTGGCATCAAACAGCAA
ATTTAAACACCATCTATCAATACATAGACCGGGTCAAAGGGAAAGAAC
TTTCTATTTCAAGCACCTTTAATATGCCCTTTGCCGAATTTGGGACAGGG
TGCTGTGTTTTCTATCTCTCCCTGCAGGTGGTCCCGAGATGACAGGCCGG
TCCTGGGCGGGAGGAGCCGACTGTGGATCCAGTTGCTTCCCAAGACAGG
CTGACAGGAGAGCAGCAAGGGCCACCCCAACCGAAACAAAGCCAGAAC
GAGCAGAAAGATGCCGTCTTCCAAAGTGGGGCTGGGAGCTTCTTCCATC
CTCCGAGGCGGTGAGGCTGCCCTGGAGCTGGCAGGAGCCACAGAGGACCC
GGCTTTGACCGCCCTCTGGGACCCACAATCAGGACCTTGAATCAGATGC
TGAGGGGCTGGACAACACCCAGGACCTTGTGCTTCCCGAAGCCGCT
GTGTCCATCAAGTCCAGATGGCAGCCGTGTCCCTACTGGAGCACGCACT
CCGTGGGGCAGGCTTTCCCTTGGGACCGATGCACCTTGAAGGCAGAGAC
GGGGCCCAATAAACGTTTCAAACCAAGTGGGTGAGGGACCCGACCGCCC
GACACCGCAGCCCGATGCAGGAGCTCCGTGCTTGGCCAGGCTCCCTTG
GGTGGTCTCTCTCTCCTCAGGGGTGGATAGGCCATCATGTGGGTGGCTC
TGGGACATCCGTCTCTCTGATTGGGTGACTTTCAGCCACAGAGATATTCC
CAGGACTACAAAGCTGGGTCCCTTGGGGCAGCTGCTGTCAAAAAGACA
AGGCCCTGACCCCAAGTAGCCAAAGTTCCCGCAGGGGTTCCTCAGGGTCTG
GTCTCCAGACTGTGCCAGCGTGTCTCCCGCCCAAGTCTGCTGACCC
GACTCTCTGTAAACATCCCGGGCCCAACCAAGCTTTACCCCAAGGCCGA
AAGCACAGCCCGCTGCACACAGATGAGGCCCAATGGCTCCCGAC
TAATCTCTGTCTGCAGTTGGCTTTGAGCTGGGGTGGGGGCAAGGCTGC
ATCTCAGGCTCCCGGAGAACTTGTGCTCCACAGCAGAGCCAGGGGCT
TGCTGACCACTTGGGCGGGTGGATCTGTCTAGAAATGCTTGAAGGTG
TCCTTGCAGGCAAGCCCGGGCGGCCCGCTTCCAGGAAGGAAGGGGACA
TTGCCAGGACTCAGGAATGAAGCAATCCAGGTTTGAATCCCGGTCCC
ACCACCTTCCACCTCTGACCTCAGGCACTTGGGCTTTACAGAGCTGCCCTT
TCTGACTCTGGGACAGGGGCTGTGAGGCGCTCTCGGTGTGTGACAGGTG
GGGGTGGCACTCTCTAACGAGGGTGGGCTGCCCAGGTGACTGACCACA
GCCCTTTCTCTCTCAAAAACGCCCGCGAGTGACCTCAGGGAGGCGAG
GGCCAGGAACCCCAACCAACCAAGATCA
Contig 105 (1820 bp)
AGTGAGCCCTGCAGGACAGTCTGCTGAGGGGTCTCTGGGCTCCTCAGAGG
CTCATGGCCACCGGCACTGGGAGATAGCAGGTGGACCCCTGCACTCCAGG
TCCAGGTCCAGGTCCAGACCCCGGACAGGCTTTCTATCTGCAGGAG
GGGGGCTCTTGGGGCAGCAGGGATGTGGCTGTGAGGCTCGTCACTCTCC
CTGTTTCTATCTCTCTCTGTATCACACACACACACACACACACACACA
CACACACACACGACGACGACACACACACAGAGGCTGACAGGGCTGCA
GACAGGGCTATGGGAGGACTGCCCGGAGTGCACCCAGATGGCCACACGG
TGGGGCTCTGTCCTCACTTTTGTGCTGATGCTTCCGCCCAGGCTGCTGG
GAGCAAGCACTAGCTTCCAGGGCTCTGACCAAGAGGGATGGGAGGGGT
CATGGCTCAACAGGCGCCAGGGAATGGGAATAGGATCTGAGGGGCGGGG
GCAAGGGGCGCAGGCGAGGCTGCACTGCCAGAGCTCCCTGCACCTGCAG
GACCAGCCACAGGCCAACAGCTGCAGGCAAGCAGGCTGCTCTGTCTCC
CAGAAGCTGGCAGACACATGGGGTCTGACAGCCCCACCCGGGCTCCC
ACAGAGGGGCGGGTCCCCAAACTCTCCCGCTCCACCTCAGAGCTCA
GCATCTCCACTGCTGAGGACGAGCCCAACACACGCGCACACACACAT
GCACGCACACACATGAATGCACCTGCAAGCACACACTCACAGTAAGCAG
GTACACACATGCATGCACACAATGAACACACATGCAGGCACACACGCATG
CACACACGACACACACTCAAACACGTACATGCAAGCACATGCTGGTCTCT
TTGTCCCGCTGAGGGGAGGATGGAGGCCAGCCCGTGGGGAGGGCATST
GGAGTGTGGGGGCTGGCTCCAACGCCCTCGCTCAACAGGCACCAACGC
TGGACTGAGATAAGCCGGGGCTGGCTCCCTTGGGGCGCTCAGCAGGT
TTGACGCCACACAGGTGGCACTGCCCTTTTCAAGACGGATGTGGCC
ATGCCACCTCACAGCCTCAGGCTCCCGCTCAGCTTTAGTGTGTCTCC
TGTCATGTACCCGGGGCTTCTCTTCCAGGGCCAAAGCGAGTTGAG
GGGACAGTGGCGCCCAATAATTACTCACCCAGGGTGTCTCTGTGG
TGGCTTGAGGCCAAGGTGCTCCATGGGGGCCACAGGCTGGCAGGCT
CACTTCTGAGAGCACCAAGGCCAGGGGGTGGCCAGGCTGGCCGCT

SUBSTITUTE SHEET (RULE 26)

FIGURE 6, CONTD.

CCCCATCTGGAATGAGGGCCTTGGCGAGAGGCGGTGCACCCCTCTTTACA
GCAGCCCCGGGGGAGAGTGAAGTCTGCGTCTATGGACCTGGGGGCTGACCT
GTACGTGTCTCGCCAGTTGCACCCCATCCATTTCCGGGTGGAAGGGAC
AAAGCCATCTTGGTCTCTCAGAGACCTCTGGAGCCTCTTGGCCCCAGC
AGCCAGCCCTCCCGGGCCCGCATCTCTGCCACCCAAAAATCACCTGT
GCCCCAGGGTCCCTTCTGGGTGTCCAGGGCGACCCAGAACTGCCCTG
CAGACACCCAGCCAGGACATGGCCGCTTGGCGGGCTGTCTGCCTG
GGGACCCCTGACTGCCACAGACAGGCGGCTTGGAGGACCATCTGCCTGAG
CCCCAAGGCACATCCACGGGGCCACACAGCCAGCGCTGTAGACGAT
GCCACTTGGGGTGGGGGAG

Contig 106 (1500 bp)

TGCCGAATAGAGGTGGAACCAAGACCCGAAAAATGTCCACATTTTCA
ATTATTAGAAATTTAGAAAAATATTTTACAGGAGTTAAAGGTATTCCAT
TCTGGGGCGGGTGGGCATGCCACGGCATGCAGGCATTCCCGACCCAGC
CACTGAAGTCTGAGCCACGGCAGTCACCATGCTGGATCCTTAACCTGCTGA
CCCCCTGGGCAACTCCAGACACTCCATATTCATGTAACTATTTTAAAC
CAAAAAATGACAAAGCTTTTCAAAACAAACACATTTTCATGGGAAGAGT
GGCATTGCTTACGCCCTGGATGGTGTCTGCGGCTTGGCGGACGACGAGGG
CCCCCGGGGAGCGCTCCGACGGCGCATCAGGACGTGGTGTCCAGGGA
AGCGGGTCACTTACGGGCTCTCGGGTGGCGCTGCGTTTCTTTTCGGC
ATCACACCCGACTCAGCACTTGGGGTCTTAAACGTGAGAGGCACTGC
GGGGCTCGAAGCCACATCACTGACCTCTCAGACTCTGTTATGTGAAAC
CCATCCGTCCACGAGACCAAGAGACAGACGAACAAACGCAAGGTGCGC
CTAGGTTGGGCACAGCATCAGGGCAGAGCGGAAACCTTGGCGAAATCCCG
GCGAAGCTTGGAGCTCGCCAGCTCTTACTTGACGCAAAACATAGGGGGATT
CAGGAAGTCTCTTTACCGCATTTGCAATTAATTTCTGCAAACTTAAAT
CGTTCCAAACCAATGCTCACTGCATGGAACAAACCCAGGGGTAGGTCTCC
CCGATCAGGATGTTTTCCCGTGGCTTGTGCGGGTGTGCCCCCTGCG
CTGGTCAGTGAGAAAGTGTCTCCACCGACGACATGAACTTCCAGGTC
CAGGCTCTCTGCTGTCTGACGAAACCTCATCTCTGTAATCTCCCGCC
AGCTCCGGGGAGGCTTCCAGGGCTGGAAGGACGGCGTCCCGTTCCAGG
GGGCAAGTGCACGCTTCCCAAAGCTCCGCGTCTGCTAGGACGCTCAGAC
GGATCAGCCCAAAACCCACGAACTGTTTCCCTCGAGGGCAGAGGCTCG
CCCTTCTCCGAGAAAGCAGCCCGACACGTCAGCAAGGGGCCAGCTGCGT
TTGTAATCAATGGCCACATAGAGTTTGTCTGAGGCGACGGGCTCTGT
CTGGGCGCACCACTGCACACGCAATATGCTGGGACACGCTCCGGGGT
CCAGGTTTATGGAATTAATAAGTTTACTGCTTCCCAAGTACATTCTTA
AGTGTAGCTGGCCSCCAGCCTGGGCGTCCGCTCCGAGGCTGCTCTCTGC
CTGGAACCTTGTCTTGGGGGACCTCTCTCCAGCCCAACCCAGCCCCG
AGCCCAAGGCAACATCTTCTTGAAGACACCCCTACCTGCCCCCTCCCG
TTCTCTCTCTGGATCCAATCTCTCCGCTTCTAAGCTCTCTTGAGGCT

Contig 107 (550 bp)

ATGGCACTCGCGGTGTGACTGAGCTACCGGACGGCGGAGCAGGGCCAC
GAGGGCGACAAGCGCGGGGTGAGAACCTGTGCGAGGGCAGGTCCCTGCG
GCTGCAGACAAGCCTCTATCGCAGGCCCCAGACAGGAGCCCCGTGTGA
CCCTCAGGCTGCCAAGCAAACTCACGGCTCTGCTGGGAAACCTCGAAC
CTGATGACTGGGTGGGTGACCCAGGACCTTGAATCCGGCTCTGCAGA
ACGCTCTGAGCCTACGGGAGTGGCCACCTCTCGGTTAGGGCCTGTGTCC
TTCCCTGGCTTCCAGCCTAGAGCAAAAGCATTAAATCACAGTGTGGCCCA
GCCCGGACCGTGCAGGACCTTAGACAAAGAGGAGGAGAGAGATGAG
GCAGAGAGGCGAGAGACAGAGGTGAGAGACAGATAGACAGAGACAGAG
GCAGAGAGAGAGACAGACAGACAGAGACAGAGGCGGAGAGACAGACAGAG
ACAGAGGTGGAGAGACAGGACAGACAGAGGCGGAGAGAGACAGACAGAG
Contig 108 (900 bp)

TTTCTAACTCTCTTACTAGTTCTAGTTTCTATTCTTTCTGGGGGGGT
TCTATATAAATTCGTTGCTGATTGGAGATGGTTTGTCTTTCTCTCT
CCAAACTGTATGCCATGTGTTCTTTTCTTGTCTTATCACACTGGCTAG
GACTTCCAGTAAACACTAGATATGAACAATGAGAGGAGAGCCAGGCTT
CTTCTCAGTCTTGAGGAAACAGTCAGTCTTCTCTATTAGAAATGAGAG
CTTTTCTTTTCTTTTCTTTTCTTTTCTTTTCTTTTCTTTTCTTTTCTTTT
AAGGAACCTCTCTTCTATCTTATTTTCTTAGAGTTGTTATTTTCTTTT
CTCTCTTTTCTAGGCTGCACCCGAGGCATATCGAGCTTCTAAGGCTGGGG
TCGAATTGGAGCTACAGTCTGATGGCTACGCCACAGCAATGTGAGATCTG
AGCCACATCTGCGACCTATACCACAGCTCACAGCAATGTGAGATGTTAA

SUBSTITUTE SHEET (RULE 26)

FIGURE 6, CONTD.

CCCACTGAACAAGGCCAGGGATTGAGCCCGCATCCTCATGGATGCCAGTC
AGTTTCGTGACCGCTGAGCCATGAAGGGAACCTCCAATAATGCACCAATT
TTAAATGAAAAAGACAAGCATCCAGCCCACAGCCTGAGTAAGGAGTTTG
GAGGCCTGACCCCTGCGTGGTCTGGGCCTGGGCCTGGGTGGTGGGGT
GGGGGGGGGTGGGGGGGACCTGTGGACCTCCCTCCTCAGCCAGGCCTG
CCCTCCTAGCTGTGCGGGGCTCGGAGGAAGCGGGTGGATGACG
GTCCCTGGGACCCCTCCTCATATGTATCTGGGTCCCTGGTCCCTCTGAGG
CCAGGTCAGGTCATGGGAGTCAAAGGTCAGCCAAGGGGTAGCCAGAG
Contig 109 (950 bp)
TAACCCACTGACCGAGGCCAGGGATCAAACCTGCAACCTCATGCTTCCTA
GTCGGTTTCGGTAACCACTGCGCCACACGGGAACCTCCTTTGCTTTTGT
TTAGGATTTACATACACGTGATAACGTGCGGTATTTATCTTTCTCATCT
GAATTATTTCACTTAGCCTAAGCCCTTCAGGGTCCATCCATGGTGTGGG
AGTGGCAGGATTGCTTTCTTTTTTTTTTTTTTTTGTGGCTGAAATCAG
TCCAGGATTATCTTTCTTTCTGTTTCTGTGGAGGACACAGGCTGCGT
CCGTGTGACGCTCTGCCGGGAATACGGGGGCCGATCGCTTTCTGAGCCAG
TGTTCTCATTTTCTTGGGAGAAGTACCGGAGTGGAACGGCTGGGTGCTC
CTGCAGTTCTGTGCTGCATTTTTGAAGACGCTCGGAGCGCTTTCCACAG
TGGCTGCACCGACTGACATTTCCACCGAAGTGCACGGATTTCCCATCTCT
TTTCCACGTTTCCCGCACTTGCTATTTTGGCCTGTGGATGTGGGCC
TCTCCGTACAGGTGTGAGGGAGTCTCCGTGCGGCCAGGCGAGGAGCGAC
CGTGAGCGTCGTTTACGTTTCTGTGGGCCACCTGCGTGGCTTCTCCGG
AAAAAGGGCTGTTTACGGCTTCTTGGCCATTCTCAGTCTGATTGTTGGG
GGGTTTGTGTTGAGTTGTGTGAGTTCCGCACGTATGGGGGCATCAACC
CTTTATCAGCTATGCGATTGGCAAGTCCGTTCTCCCATGTTCCCGCGGCC
GCCTTGGCACGTGTGGGGCGTCTCTTGGCTCTTCTTGGTGCAGAAGGC
TTCGGTCTGATGTGGGCCCATTTGTTATCTTCTTTCTTCTTCTCACCGT
TGTTTTGATGTGAGATGCAAAATCCATTGCCAGGGTCTGTGCCGAGAAC
Contig 110 (306 bp)
CGCCACCTCAATCGCCGTTTGTCTGCAACACGGTCCAGATAACCAAGCG
CACCTAACAGGTCGAACACTGCCAGAACGCGAACAGCGGCTGAAGCCG
ATGGTGTGAGCCAGTGACCCGACACAGCGCAACAGCGTACTTGCCAG
CCATGCGGACATCCCGTTAAACCGTTTGGCGTTGCCACTTCGTTACGAC
CAACACATCGGAAGAGAGCGTATCAGCGCGCCAGACAGTGCCTGGTGG
GCAAAACCAACGATACACAGCAGCATAATTGCGACATACGGGTGGTGAA
CAGGCC
Contig 111 (800 bp)
GTTTTCCATGATGACACAGGGGGGCGGGACCGCAGCAGGGAAGGCTCCA
TCCTGGCTCTGTAAGACCTTGAAACACCTCATTCCTCTGCTTGGCCT
GCTCTTCGGTACGCCAAGTTGCTGAGACTGATGTGGGGATCAGTGGGGAG
CAGGAATCTTTCTGATTACGCCGTTTCAAAGTGTCCCAAGCAGAAGCTGT
GATGGCAATGCCAAGGCTATCCATGGAGGTGGCTGTGCCAGGGGCCCAT
TTCTTGGGAGCCCATTCAGGAAAGGAATCTTGTAGCCCAAGGCTCCAGC
AGCCAGTGCACGGCCCTGGGACTATCCGGGTAGATCAGAGGGAGGAACA
GAGCTGTGGATGGTAAGCAGGTGGCCCAAGTCCAATTTATGTCTGTGGTC
CCAGCAGGGTGCCAGGAGGCCCTCGTAACCTCTTAAGAATCTTGGTCTG
GTCAGCTAAATGTATGACCATTGTACTGAGCACACATCCCGTTAAGTA
GAATTTTCAAGGATGACTAGGAGTTTGCCACCTGAAGGCAGGAAGGGCAT
TCCAGGCAGAGGTTACAGAGGTGAGAGGGAGGCTGTGACACTTTGGGCGT
GCAGGGGGTTTGTGACTGCAGCTGGCACACAGTGTATGCCAGGCCCT
GGCAGGGCTGTGTTGGTGTGAGAGGAAGGGAGAGGTGAGTTGAGCCC
AAGGTCTTCCAGGCCAAAAGACTGAAGGTGACCGGGCTGTCCGGGGCTG
GCCCCAGACCAAGGAGGAGCAGGTGGGAGCTGGCTCTTGTCCGGGGAC
Contig 112 (3062 bp)
CACACCCAGGAGAGGAAAGACCCACACAGTCTGATGACAGCTTGGCTC
GGGGCTGGAGCCCCGAGTTATAAATGTCCATCACGAGCTGTGTTCTGTCA
GAGCCATCAGTGGGAAGGCCAGGCCAGCTCAGCAGCCAAAAATGAAGAG
CTAGGTCTGGGATTGGGCCAAGCAGAGGGCACAGGAAAGCCACATAAAC
AAGGCACCCAAACCCCTGTATCCACCAATGTACATTTCAGGTACACCC
CCTGGTCTTCCGGGGAGGTCCCTAAGATCCGGTGGCAGGGGGAGGAAAA
GTCTGACTGGATTCTTGACAGGTGTATCAGCGGAAGGCCAGGAGGAGTG
CTCGGGCACTGCCACCTCCAGGGGATGATGGTCTATGGACAGATGGCA
GTTATGGGAGGAACCTCCCCGTGGTCAGAGCTCTGGGTGCTGTACCTGG
TCATGCATTTGAGTGGAAAGGAAAGAAACATACAACCTCAACCCCAAGC

Contig 113 (1300 bp)

AAACCGGATAAATACAGGTGACCCACAGGCAGAGCTGAAGTACAAACAGT
TCACAACGACGCCACCCAAAATAACGGAAGGCTCAAGGCTAAATCTGACCC
AGATGAAAGGCCCTTCTCACGGAATAATGGCAAAGTGGCGCTGAGAGGCATG
AGAGGTTCTGAATAGATGAGGAGGCTCCGCGTTTTCGCGGCTCCGAGGA'T
CAGTGACGTCACAGCGCCAAATCTCTCTGAAACGCGCTCTGTAGTTTCAGTG
CAGCGGACAGCCACTGGCAGCGCCCTCGCTGCAGAGACAGCCCCAGCTGG
GTCCTTGAGGTTCTCTACAGCGAAGCAAGGGTCTAGAAAACAGCAGCTCT
CTGGAAGGAGGAGAAGCAGCCGATGGATTGGCATACGGCGACAGAGAA'TTC
CTCGGACAGTGGCCACAGGAGAGGGGCTGGACAGAGACTGGTGCACCCGAG
CGGGCGCAGGAATAAGTCCACACCCACAGCTACCATCTCGTTGTTTA'TT
ATT'TTTTCTTTTTCAGGGCCACTCTGGGGCATGTGAGAGGCTCCCCAGCC

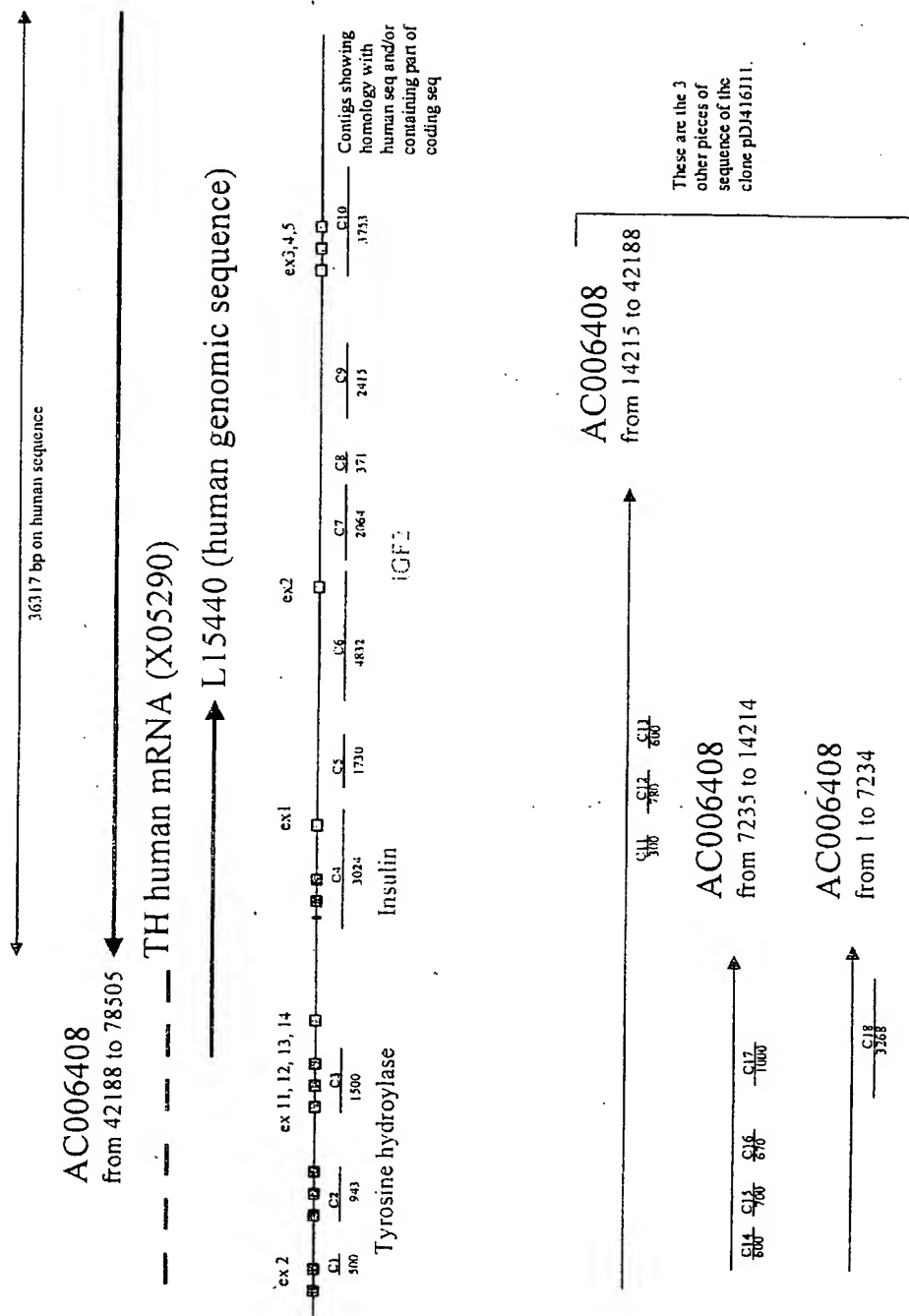
FIGURE 6, CONTD.

AGGAGTCGAATCGGAGCTGCAGCTACAAGCCTACCCACAGCCACAGCGA
CACAGGATCTGAGCCATGTCTGCAGCCTACACCACAGCTCCCGGCAATAT
TGGATCCTTAACCCACTGAGCAAGGCCAGGGACTGAACCCACGTGCTCAT
GGATACTAGTTGGGTTTGTACCACTGAGTCACAGTGGGAACCTCTTTAA
TTTAATTTTGAAGGTTCAAGACTCTTTAATTTTACTGAGGTATAGA
TTATATTACGCACCATTCTTTCTGACTTCGGTGCACGGCTTTTCAACAA
ATGGGTGCTGGACCTGCTGGGTGCTTCTTCAAATGAACCAAGCCCTC
CTCGCGCGGTATGCAAAATTTAACTCGAGGGGCTCATAGACATAAAGCT
AAACTCTAAAGCTATAAAATTTCCAGAAGAAAACGTAAGGAAAACCTTG
GGGTCTTGGGCAAGATTCTTACCCATGACAGCAAAATACAATCTACA
GAAGAACTGGTGGCTTTATCGGCATTTAAACACCTGCCCTTGAATGA
TGCTGTCGCAAAACCGAACATGCAGCAAAACGGATGCAACTAGCAGGTCT
CACACTCAGTGACCCACCTCAGAAAGGGAAGACAGCCACGTGACATCC
CTTAGATCGCAAGATGTAAGACAGGCCCGGTGAACCGACCTCAAGAGAG
AGACAGACCTACAGACGAGCAATTTGGGGTTGCCGAGGGGATGCCGG
Contig 114 (3000 bp)
TGTCAGACCCCTTGGCGGGCCAGGACCCCAAGGTGACCGAAGGCCCTCA
GGCCCCCAGCGCCCCATCCCCCTCTTCCGACACAGGATTTTTTCC
CACCAGCTCTGTTCCTTGGTACGCTCTCACTTGAGCAGCCTCAGGGT
CTCCCGGTGCTTATCCACGACAGCGTGACCTTCTTGGTGTGCAACCC
AGGACCCACGCTGGCCAGCCACGCTTCCAGAGCACCCCGCCATCC
TCAGAGTCCAGAGGAAAGGCCCTTACACCCAGAAACCAAAACGCGAGA
GACTCTGGGACGCCAGCAAGACGTACACTGACTCCACCTGCTTCAGGC
ACCGAGSCAGGGGTGGGTATGAGCGACCCCGTGAAGGGCTTCTGTGTC
CATCGAGGGGCTTCCAGGGGCTCTAGACGGGATGAGTGTGGCAACATG
TCGCCGCTTACAAAGACCTCCAGTGCTGCTGGGATGGGTCCCCCGGC
TAGAAAGCAAAGGATTCCAGCCAGTCCAGTAGGAGGCGGCTCGGAGG
CTGACAGAGCGCGGGGGGCTGACACCACTTCGCAAGCCCGGTGTTC
AGGGGACGCCCGCGGCTGCAGCGGTGCGCTCCGGATAAGCTCCTA
AGAGGCGCGCTTCCCATGCACGCGGTGCACACACTCGCTGCCCGAGGG
TCCTTCAGCACAGACTTCTGGGACGAGGACCTGGCAGGGGTGTGGCT
CTGGGGAAGGGGTCTGTCCAGGAACCTGTCTGCTTTGGGGTGGGC
GTGGATATCCCGTCCCAACCTACAGAAGGAGGGGCTTAAAGAGGCC
TTTGGTGTGAGGGGCGCAATCTTTGGCTTTTCTTGGCCACTTGA
GCTTGACGTCTGGTCAGTGACTGGAGCCAGGGCCAGAGGGGGCAGCGG
GGCTGAGCGAGGTTACAGGCAACCTCTCTCGGCCACACTCCCGAGGTG
GGCAGCTACGGGGCCCCAGAGACACAAGCCCCAGGGGTCTTCCCCC
GCCCTTCCCCAGATCACAGGAGACCAAGCAGCTCTGCTCCCCGCTG
CCTGAGAAATGCCCATCTGGGTACCAATCACCTCCAGAGGTAGA
CTGGGGGGCCAGGACAGGGGGACCCAGTTACAGAGCCCCAGGCGAGCT
TCCAGGGGGCAGGGGACTCCGTTTGGGGCACAGACGAGGACAGAGCGG
CTGATGGATTCTCCCCGCTTACGGGATGCTGGCTGCTGGCTCCAGGA
GCCGGCGGTGCCATCTGATCTGATTAGGCTTGCAGTCCAGCTGGGGG
GCACAGCTCGGGGCTCGGGGGCAGGGAAGAAGCGCTGTGCGCCAGC
CGGTACGGCTCGCTTCTCTTCTTCTCTCATTAAAGTGTGAGAAC
CATTTATTGATTTTTTAAATCAGGACGTGCTGTCCGTGACACAGCAAGT
GAACAAATCAGAGCAAGAGAGGCCAGGGCTGAAGCCCCAGAGGGCGG
GCCTCCAATCCGGTTGTGCCCGGGCTCCAAGCCCCCTTCTTCTCTGG
GGTCTTGGGCGTAGTGGCCAGGGCAGAAATGCACCTGCCGTATCTTGGGA
GGCTTGGCCATCGCTGGCTTCTGTCTATGACGCACCGTCTGTTCCATATC
TAGGAAACAGCTTCCGATTAAACAGGCAGGGGAGGCGGTGTTCTCTCT
TATCTGCCACCATCGGGCTGGGGCCAGTGGAGCCAGCCGCTGACT
TCCCGCTCGCACGCAAGGCACTGATGCAAGAACAGGACATCCAGCCCC
CGCTCTCAATGCCCGGGTCTGAGAGCATTCGCCCAACAGGCTTGGG
TGGGACAAGGGATGGAGCTGTGCGCCAGGGGCTGGCTGGGGCAGAGGG
GGCTTCCCGTGTCTGCCCGTGGCTCCAGCACCTCGGCTGCCAGGCTG
CTCTGGAGAGGTGCCCGGGGGCAGGGGCCAGGGGACCTGTTCTGCC
CACGTCTCTGTCTGCTGAAAGTTCCACAGACCGTGTATACCTTG
GGAGTCAGGAGGATGGGGATAGTTGGGGCTTGACGTCTGTTCTGAAAA
AACACCGTTTTCCCTGAAATATATATGATTAATTTTCGTCAAGATAAA
ACTGTGTATAGTTTTCTGTGATGAGAAACGATCCATCTTCTTAGAAA
GCCGTAAGAGGTACAGGAGCTATAAGGACAAAGATGACAGATGCCCTTA
ACGCACACCAATGTGCGGTGGCCCCAGGGGACCGCATAGACGGGGCGG
CTCCAGATGCCACCGTGTGCGAGGACACGGTTTCAGGGTGGCAGAGTAT

FIGURE 6, CONTD.

TCCTGGGGGGGGGGCTCAGCGGTTCCCATTTCCCCCTCCCTTCCTTCC
TTCATTTCTTTCTTTCTTTCTTTTGTGGTTTATAGGGCCGCACCCG
CGGCGTGTGGAGTTCCACGCTAGGGGTCTAATCAGAGCTACAGCTGCC
GGCCTCCACCACAGCTCACGGCAACGCCGATCCTTAACCCACGGAGCGA
GACCAGGGATGGAACCTGGGACCTCATGGATCTTAGTTGGGTTGTTCCT
GCTGAGCCACAACGGGAACCTCAGCCATTCCTATTCTTGCTCCAGTTCC
AAGAATTCCAATTCCTATTCTGTCTTTAAGGCCAGAGCGACAGCCAC
GCCGAGTCCAGAGCAGGGCTCAAGGATGCTGCTGTTGACTGTGTCCGT
GGGCGGGGGAGTTGATAAGAACCCCAACACAGGGTGGTGGCCAGCAAC
GGGGGAGGGAGGGGGGGCTGTTGGGAAAAGTCCCTGAAACCCATGG
GCTGCCCCCTCCAGGCTGGGGCAGACCCCGAGCCCATGGCCCGAGGAG
AAACGGTCCAGCCCGAGGCTGGGCTCCCGCACCCCTGCCCTGACCCCGC
Contig 115 (1895 bp)
TCATGGAAGCCCTTATCACAACCTCGGATCCAAAACCCACTGCGCGAGTC
CAGGGATAGAAGTTCGCATCCCCACAGACCTTATGTTGGGGTCTTAACCCAG
CTGAGCCCATGGAACTGGGTAACTATTTTATAGATGTTCTTAGGGTTT
TTGGCCTTGCCCTGACGTGGGGACCGTCTGGGCCAGGGATCAAAACCCG
GCCACAGCTGTGACCAAGCAGAGCAGTGACAGCACCGGATCCTTAAGCA
CGAGGCCAGCAGGGAGCCCTGTGTTTAGATTTTGGTGGAGATACTGCGT
GGGATTGAGGATATTCATTTGGGGCTGTTGGAATTGCCCGTCCGTGTTT
AAGCAAAGAGAAATCCCTTCACTCTGTGAACCTGTTGGGAAATCCTTAG
TCTCTTGAACCATTCGCTGTGTTTAAGAGTGGTAACCTGCGCACCATAA
ATGCCAGACCCAGCGCTTCTGAGATCCGCTTTTGTGCAAAATATCTCG
TTTGAATGTTTGTATCCCCCGACACAGACAGGGTGGGCGGACCGCGCG
GGGACCCGACGTGACCATCGTCTTCTGTATCCGCCCTTCTCCGGCAGC
CGCCCCCTGGTTGCCCTCTGGCTGCTTTTAGTGGAGGAAGTGAAGCTCCG
CACCCAGACCCCGAGACCGCAGGACCCACAATGCTTCAAACACCTGCCCT
CTGACTTTTACAGGTCAAGTTCCGCCAACGCCGAATTTGCACCGATTTGGCT
ACAGAGAGCACGGTGGCGCCAAGCCTCCACTTGGAGTTTATAAGGTCTC
CCTCCAGCTCGCAATGAAATGAGCTGTGATAAGGCAAAGACAAATTAG
TATGAAATCCAGATGCTTCATCTACAATACAATGACCCGCGGATTTGGGT
CTGAGCGACTGAAATCAAGGTGGGCTTCCGGAGGGAGGCTGTAGAGGAA
AGGCATTCACCGAGCTCAGGTCCGAGACGCTTCCACACCCCTAAGAGGG
CTGAGACGGCAAGTAGGGACCAAGCCCCGAGTCCGGAGAGCTGGGACG
AAGGAATCTGAGGTACCCCCACCTGGGGAGGAACTGCCTAGAGAAGCG
GGGGCGGGAAGCAGGGGATGCCAGTCCCAAGACAGGGACAGGGCGGAAA
GGGCTCTCTGACGGCCCTCAATGCTGCCACAGTGTCTCGTAAGAGGGAG
CCAGAGAGAATTGACACCGGGGAGACCACGGGACCACGGAGGTGGAGACC
GGGCTGCCCCGCGTGCAGTTGCTCCCGAAGCCGGCCCTCCCCAGAG
CCTTTGGGAAGAGGGCGCAACCTGCAGTTCTGCTACTCGGGACAGGGAC
AGGGACAGCCCCCTGGAGCCGCTCTTAGGGGCAGCATCCCCAGAACCT
TCCTTAACAGACCATCTGGAGAGAGATGGGTCTGGGCTGCAGCTCCTGGA
ACTGTTTGGCCACCCGGCGAGCAGTGGGTGCCAGCCTGGGCTGCC
AGCCTCAGGGCCGGGGAGGGCTGAGGGCACTGGGGCCCGGCTCTGGGACT
CCCCTGCTCTCTGCCCGTGCAGGACAGCCACCTCCAGCATCTGCTTCT
GCCACCCACATCCCCAGGACCGTCAGCCAGGCATGCCCTGGGCTCGGC
CACTCACACCACAGGCCAGGAACCAAGGGGGCAACACAGAAGGGCAGTT
GCCATCTGCAGATGGAATGCACAACTGGGGTCCGTGATGATGGCAGGCT
CTGGGCGCCCGGGTGGCAGGGGAGCCAGGACTGTGCGCCATCACAGGA
AGGGCATGACGGGTGAAAGCAAGAGTGGAAACCTTGCCACCCGCTGG
CGGCACATACCGGCCACCTGCAGCCCCACCCCATTTGTTTGTCT

FIGURE 7



SUBSTITUTE SHEET (RULE 26)

FIGURE 8

Contig 1 (1040 bp)

GCGCGCCGGATCCTTAATTAGTCTGAGAGATCTGCGGCCCGGCCAGGGTCTGCTTCTG
GCCAAGTGTGGGGCTCTGCTCCATCTGGCTCGGAGGTCCACCCATGGCAAAGCCTGGGG
TCCTCCCACTGAATATTTGGGGTCCACTCGTCCAAAGGCTGGGTGTCCAGTGTGCCAA
CGGTACATGGAAGCAATGTCTTCCCAAGGACCGTCCPAGGTGTGGTCAGGCCTGGACAGC
TGTGAGTCCCTTCGGGACTAGACTTGGTGGCCGAACCTAGGGACCGTGCCTGAGGGCCC
CCACGAGGCCAGGTGTTTGGCCAGGACAGAACGSCCAAGGGTGGCCGAGGGTCTTTT
TGTTTGTTTTTCTTCTTTCTTTTCTTTTCTTTGGCCGAGGGTCTTAAAGCGCTCTCTCTG
CTCTTTGTCCCGATCCTGAGCGGGCAGTGTCTGGTGGGTGGGTGCTGGCCAGCCGACG
CAGGGCTGAGAGAGCCCGCTTGTCTACTAGGGCGCCUGGTGAGCCAGCCGCGCATGCCG
TGTCAGACGTTGGATGGGGCAGCGAGGGGACTGGGGTGCCTTACGCCCGCTGGGAACCC
CGCCCTGTGGAAGCCGCTGTGCTGCCACAAACAGCACCGTCCGACTAGCTGGTGAATCAG
CGCCCGTGCCTCGCGTAAATCCAGGCGCTTCTGCTTAACTGAGCCCTGACCCACACCC
CCTTGGCAGCCGCTCCGTGGACCTGGGGCGATGAGGTGAACCGTGGGCTTGGCCATCGTG
GTGGCAGACGCTGGCACACCCGTGCGCTTCTGGGCCCTCCATCCAGGAGCAGAGTG:
GCACCCAGTGGGGCTGGGACAGGAGCCGCTCCACCTCCGCCCTGAGGGGACGGGACTC
TTTCGACCCGGAGTGGGAAGGACATATGCGGACGATGCCAGACCTGTCTGTGGGGGA
GGGGGAGAGGCCCTCTTTGGAGAATCCAGGACGGGTGAGGAACGTGTGCTGGACCGGC
CGGGTCCGAGCTGGGCTTG

Contig 2 (9234 bp)

GGCAACCAGGGGAAGATGGGAAGCGGGTGCAGGGGCGTTTGGCGGGCCAAAGGACCAC
CTTGGAAATCTGGAGCCTGGCAGGAGCGCCGAGGGTTGAGGGGCTGGCTTGGGCAGGGC
TGGCTGGCAGCTGGGAGCCTGGCGGGTGGAGTCCGGGCTCCAGGTGCCCTATAGGCA
GGGCAACATCGGCATGGGGGTGACAGGCCCGAGCTGGGGTGGCGAGGGAAGAGGGGGGA
GACAGGCATTATCCCGGTCAATTTGGTTTCAGGTCTGGCGGCTGGTGGTCAGGGGA
GTTGGAGAGAGGTTCCGCCCCGGGGCTCGGGCAGCGAGGTGTAGCTGGCAGCTGTGGGC
AGGTGAGGACAGCCGTCTCCCGGGCCAGGTGAGTCCCTTCCCTTCCCGAGGCCTTGTTC
TCTGGCTCTCTGATCCGAGGTTCTGGGGAGCGAGGGCCGGCGAGGCGAAGCGGCTGAC
CCCCGGCAGAGTGGCGCGGACGACAGGCAAGGCGGGCAGAACAGGTGACACGTCTCAG
GGGAGCTGGGACCGGGCGGGGCTGGGGGCGGGGCGCTCCAGGTGGAAAGAGCATCT
CAAGCGAGTCTGCTGGCAGACGAGGCGAGGCTGCCAGCAGGGAGGACGCAACAGCGG
GGGGATTCCAGGCCCGGGTCCGACAGGACCCGTCGGGGGTGTGAGGACAGTGGGGTCCC
CAGCCGCCACTTCAACCACTGCAATTCATTTAGTAGCAGGTACAGGAGCGGCTCTGGCCG
GGCTCTTGAGGCTGAGCTGGAGCTTCAGGGCCGGAGAATCGGAAAGAGGTGCAGTG
TGCCAGACAGACGTCACTGGAGGGAGCAGGCGCTGGGGACGGGCCCCAGAGAGATTTC
GGCAGCAGGGAGGCTGGCGGGGCGCAGCTGCGGACGTGCGTTCCACGCGAGCACTGCGG
CCAGGGGCTGGCGGGCAGGGCCCCCGGTGTCTTGGTGGCACTGTGCGCCCTCGCCGC
TCGCCCTTGGACTGGCAGGCGAGACAGGACAGCAGGAGGTCAAGGGCACTGACG
AGACCAGACTAGCGAGGGGGTGGGTGGAATGGATGTGACCTCTGGGGGAGGGAGGT
GGGGACGAGGCGAGGGGCGAGGCGCGGAGCCTGGCGGCGAGCGAGGCAAGGCGGGCT
CTGCGGCTGACAACTGAGCACATATGGGTACCTTTGCGCTCGCACCGGAGACAGTGAGT
GTCTGGCCCCGGCTGCGGCCCTCCCGGCCCGCCACTGCCTTGCCTTCCCCCTCGACC
AGGGCCCTCTGCTTCCCAACAGCCTGCTTCCAGTGGGGGTGGACACACTGCCAGACCA
CAGGGCGGAGCGCAGGATGTGCTTGGAGGACATGACACAGTCCGGTGTGACGGAGAGGG
ACAGACGTGACCGCGTCCGGCTTCTGGTGGCGCAGGTCCAGGGCTTGGCCCCCAGGC
CAGCGCCCCACCCCCACCCCTCATGGCGCTTCTGTGTCGCGAGAACACTCTCGGCTG
GCCCCGCGGGGAGTGGCACACCCAGCGTCTGTCTTCTTGCCTTCTGAAGGAGCAGT
GCATGACTGCTGCTCTTGGACCCAGAACCTCAAACGACAGGTGAGGACAGTCCCGC
CTGCCCCACACGTGGAAGGGGCTGGGCGAGAGCGGGCGCTCACGGTGGCCCCCTCCC
CCTGACAGATGCTCTACCCAGCTCATGCTGGGCTTGGACCCGAGTCTTCAAGTC
CTCTAGCTCTGACTCAAGAAATGCTGCATTCTGGAGCCACTACACTACTTGACTCAGG

SUBSTITUTE SHEET (RULE 26)

FIGURE 8, CONTD.

AAGAGCAACGCTCTGAGCTAGCTCCACGCGTGGGTCCATCTCGGCCAGGTTTAAATGAGCC
ACTTTCAGGCAGGGATTGCACAGGAGGAGGGTGGGAAGTGGCTCTGCTCAGACCCCTGA
ACAGGCTCTGAGATTCTCCAAGGGCACAAAGAACGGACGATGCCCTGGGGTCAGCGA
CAATGCTCCCTGAGAAATCTTGGCACACAGGGCTGGGCTGCGAGGTGGCCCTCGCCCC
ACCCACAGCCTCCTGGAGGACACCGTCCCTGCTCCAGAGCTGGGGGGCGCCACAGCT
GGGGCACAGGGAGCATGGGCCCGATTCCAGGCTGGGCTCCCTCTCGTGTCCAGGATCTC
CCCGTGTCTGTCTCAACAAGCCCTGACTTGGAGGCCAGGGTGACCCCTTAAAGGGG
GAACAGAAGTTCTAGAGGAGCCTGGCCAGCTTTGGCTTCCCTAGGGCTGTGGTGACCA
CCTGGGCCACGGCCAGGCCACCCACCCGCTCTTCCCTGGGCCCTCCCTTCCCTTCCCT
CGCACCTCTCCCTGGCTGCACCTGGTGACACGCTGGCTCCAGCCAGGGCTGAGGGGG
ACCAAGCGGGGCCCTTCTTGAAGCCACCTGCAGGCCGGCTTCTTGGGAAGGGGCTGCT
TCTTCCCGGCCACCCGCGGGGGCTTCTTGAAGCGGTCACTGGATATTTTGT
CCTTGTGACGCGGAGCTTGCATAAAGCAGACACTGAUCTCTTGTCTCCGGGAGCAG
CGCTCCATCACCAACACCTGGCCGGACACAGGCGGGCAGCCGCTGGGGGAGCAGCG
CGGGCTCGGGCCGGACAGCAACGATCACGCGCGGAGCGAGGGCCCGCGGCTTC
TGCAGGCCCGCCACCTGCCAGGCCAGCGGTGCCATCTTGCAGGCTGGGAGGAGCG
TGTGGGGCAGAGCTGAGAAGGGGCGAGGCACTGGGGGGGACAGCCGTGTTCCACA
CTTTGCAGAAACCTTGGCCGGCTGGATGTCTTGTGGAGAGCTGGGGGAGGGGACAG
GCAGGAAGCCGTTCCCGGAGCGGGTAGGAAGAGCCCTCGGCCCTGGGAGGAGGAGGA
GGGAGGGCAGTGAGATGGAAGAGCACAGGGGCTCGAGGCTTCTTCTGGAACAGGA
CTAGAGAGGAGGAGCGGGCAGCTGCTTGGGATGCTTGGAACAGGCGGCCAGTGTCT
ACAGGAGCTGACTGCGCGCGGTCCCGGGCCAGGCGGGCTGGGAGGGCGCTGTGGT
GTCAGCGCCACTCAGAGCCCTGGCAGAGGGGGCTGGGACGGCTGCAGGACAGAGCTC
AGGACACAGATGGGGGCGAGGACTGAGTGGGGCAACACAGATGCTCCAGGAGGTGGCCA
AGGAGTGGCCTTGGGATCCAGGATGGCCCTGGTCCAGAGATGCGGCAGCCCAAGGGA
CCAGGCCAGGGCCGCGAGGGGCCACAATCTGAGCAGGGCTCAGGCCAGGGCAGAGGCC
CTCCCAACCGAGCCCTCCCTGGGCCGCTCTCC
GTGACGGCAGTGGGCTCAGATGGGGCAGACATGAGACCAAGTCCAGGGAGAAGCGGGCC
CCTTGGCTTCAATCAGGTGGCTTTCAGACCGCGTCCCGTGGCTGGCAAGGCCACAGCGC
TCAGGAGCACACAGACCCACCGGCTCCCCAGGTGGGGCGGTGACATCAGCCCTG
TGTCACACAGCAGGAGCTGGCAGCTCCCGACCGGGCTTAGGGAGCGGGGACCTGAGCCA
CCTTCCACCGCCCAACCCACCGTGGCCACACAGGGCCCGCTGCTCTGGGTCTGGGG
CCAGGGCCCGCAGGGCCCTGGCACTGTCTGCCCTCCCGCTGGCTCTCCGTCTCCAGTG
TCCCGCCAGAGAGCATGGGGCCACAGGCTGAAATGCCACCTCTTCTCCCTCTGAGG
GGGCTGAGGTTTGGGGTTTACAGAGTGGCTCCGGGTGGGTCCAGGCCACGCGAGG
CAAAGCGGACCCAGGGAGTCCCGGGAATGTGGGACAGCCCGCTAGATCTCGGGGG
GGCCAGGCTCTGGTGACCTCCATCTGGGGCTGTGGGCTTTGGTCAGTGGGGAGGGTC
ATGACACCCAGCCACAGCTGGTGACAGCCCTGGACGTGCCGGCTCAGGGCTGGCCTGC
CCCTGCAGCTTGAACCCCTGTCTCTGGGAGTGGGGCGCAGGGGGCCCGGGCCAGGG
TGAGAGACGAGAGCTCTCTTCCAGAACTTCTGCTGCGATGAGGACCCAGCAGGGGCC
TCTCTTACAGAGGGCTCTGCCGGCTCAGGGCCCGCAGAGGGCCAGAGGCTGGAGG
CCGGGCTTGGGAAGAGGCGGACTTCCAGAAACAGCTGCCGCTCCGAGCAGCCAGC
GCCACTTGGGAGGGGGCGCGCCCCCTGCCCCGCGGGTCCACTGCTGGGGCCGCCA
CAATAAAGTTTGTCCCTGCTGGTTACTGTCCGTGTCTGAGAGGTTTCTGGAGCCTGGCCA
CAATGGGCGTCAGGATGCGGCTGGGAGGAGCCTCGCGAGTCAGAGTGTCTGTGCTCTCG
ACAGGCCCGGGCGCCCCAGCCGCTGTCTGTGGACAGATGGGTGGGTGGGTGTCTCG
GAGGGGTTGGAGAGGTTGGCGGGACGAGGGGCTTCTGCACTCTGTCCAGGGAAGCG
GGGACCAAGGAGGGACAGCCCGGTCAACAGGAGGCTCTGTCCCTCTACCCCGCG
GACAGGTGAGCTCCCGGAGCGCCCTTCTGGGACAGGACCCAGGCCAGGCCACGGCC
CCCCCACCCGTGGTCCCTCCCTCCACCGCCCGGCTGGGGGGCCAGGGGCCAGGGCC
CCCGCTCCCGTTGGCCCTCCGAGGTTAAACGACCTCGCTTGGGAGCTGGGGCAGAGGGC
AGGCGCCAAGAGTGACCCCTGGGACAGTGGCTGTTTGCAGTTCTGAGGCGAGCCGAGA
TAAAGCGGCTGTTTCCAGTGGGCTCAGGGCCAGAGGGGGCGAGGGGAGCCCCAGTC
AAGGCCGGGCGCTGCTCGGGCTCCCTCTGTGGGAGGAGGGGGCGGTTGCACAGC
AGCCCTTCCCGCCGCGCCCGCGCGCAGGCACCGTGGGACCCGGCTTGTGCCCT
CCCCCGCCCTGCTCAGGGGCCAGCCCTCTTGGTTCCAGGAGCCCGCCCGCCGAGG
CGGCCAGAGTCCAGAGTGTAGCCCTCCACGTGTGGGATCTGTATATGCGACAGC
TTAACTCAGGCCGAATTTATGGGTCTGGATTGGGTGGGACGGCCCTGCACAGCGG
GGCTGGAAGCCTAAGCGGTGGGCGTGGGGTGGAGGCCCCGAGACACAGGAGGAGG
CTGGGACACTTCAAGGCTGACATGCTATGCTGTACGGATAAATGC

Contig 3 (5347 bp)

AGATGTGTATAAGACAGGGGCTGGGTGGGAAGGACAGAGGTTGGGGCGGAGGAAATG

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FIGURE 8, CONTD.

GGATGCAGAGCCACCGTGACGGCTCTGCTGGCCTTTGAGCCTCGCTGAGTCCGAAGAAG
CCCTCGGGCTGGAAACAGACCCCGGCCCCACCCCGGCCCGGATTACCCC
GGCATGGCTGGAGGGCCCGAGAAAGCCACCCAGGCTTCCCGTGCCGAGCTGGGTGCTGGGC
CCAGCCGAGCGGCTTGACGCCACGCTTAGCCCTCCCGAGGAGCCAGGGTCGGAAGGA
AGAGGCCGGCGGAGGGCGGTGGCCGCTCAGGCTGGAGGGGGCCCCGGGTGAGGATGGG
CCCCAGACGTCCCGCTCCCGGCCATCCGTACGGAGCTGTACCCAGGAACGTGCTCC
AGACGTGCTTCTCCGCGGAGGCCCGAGCAGGCTCCAGGCGCCCCACCCCGAACG
CCCACGCACACCTCGGTCTGCGAACACCTGCCGTATCCGGTGGCCCCGGTCCCGCC
GCCCGGCCATCCGGGTGCCCTTCTCCCTGGGTGGGGGCCATGCCCTCAGCGGGCAC
CGAGGCTGTGCAGGTCTGTCTGACTCTTCCCAAAGACGCGAGCGGGTGGGGCGCC
CCGACCTCGTCTGAGGCCGTTTGTCTACTGCTGTCTCAGAAAGGGGTGCCACGGG
AAGCGCTGTTCCTTGGGCCGCAAGGCAAGGAGGCCACCCCAAGTGGCTGAGGGCAAA
TGGCCCGAGGCTCTAAGGAGTCCCTGGGGGCCGGCGGCCCTGCAGCTTGAGGAGGAGA
GCTTGGCTCTGCTCCCGGGCAGGTGAGCCACGGCAGGGGGCTCCCGAGCAGCTTG
GCAGGAAGCAGTGAAGAGGGGTGAGGATGAAGGCAAGGGGGCTCCCGGGACTTGGGCA
AAGCCCTGAAGAACTGAGTCTCGGAAAGCGCGAGCCCTCAGCCGAGCCTCGGCCCTC
CGAGCGATGGAGGGCGGCCACCTGC3GCCCGAGGTGCAGCTGTGCATCCCTCCCTCG
GGCTCCCGCTGCCCGGCCACCACTCTCCCTTTTGGCTTTGATCACTTGAAGT
GCGACAGCTTGTGGCGCTGAGCCCCAGAGACCGTGGCCCCCTGCCCGAGCCCCAGGG
GAGCGTCCACCTGGGCTGGCTGGGCACTCATCCCTCCCGGATGAGGCCCTTCTAGCCT
GGCGCGCCCCGGGAGCGGCAGACCCAGCCCTCGCCCCCTCCCGAGTGAAGGTGCTGC
CTGTTGGTCTGGGGAAGCCCTGGAACAGGGGGCGCAGGTCCACACGGGTGCTCTGGCC
TCCAGCTGCCAGGGAGGGCGCGCTCAGGCCAGGGTCCCTCCACAGAACCCCGAGGGC
CTTGGGAAACCTGTCTGTCTAACAGGGCGCTCCCGGGACTCCACGGAGAGGTGCG
AGGGACCCCTGAGCACCCACCCCACTAAGGGGGCCAGCCAGCTCGCGGTGACAGGAGC
CGGTGCGCGCTCAGTGCATACTGCTCTGTGGCTTTGTGTGCGCTGGGTGGGGT
AGCGGAGGTGCCGAAGGGCGAAGAGCCACCCCTCCACTCGGGGACCTATTTCAGCAAGA
AGACGGATGGGACTCCCGGCATGGCAAGGAACAGGATGAACCTTCTGGAACGCACA
GGCTTCCACGGCTGACCGGTCTAGGAAGGGCGCTCTTAGGCCAACTCCACCTCCACCG
TCCATTCCCGAGCCCTCGAGAGGGGGCAGATGAGCCGCTGCAGCGTGAGAGAGCTCTGG
CGCGCTCCACAGGGCAAAGTCCAGGGCACTGACCTCAGAGCCCAACAGGCCACCGGG
GCTGGGCCACAGGGAGCCGGGGCCAGGGTCAAGGTCAGGGCCAGAGTGGGGAAAGG
GTGGCTGTGTGCTTGGGGCGCGGGCGCGCAGAGCGGCCCTCGACCCCCGACAGCCCT
GGAGCTGAGTGAAGCCCGCGGTACCTTGGCTGGGTGGGGTCTCTGCGACCCGCGAC
CCCAGCTCAGGTATCTTGTGTACCGCAGAGGGGAGGGCTTCTGAGCAGGGACAGGG
TGGGCGCGCAGGAAGCCCCCTCTCTGTGAGGTGCCCGGCCCTGGAGCCTCTCTGGG
GCATGCCACCCCTCTCACAGACGCTTCCAGGAGCCCCACTTCTGTGCTGGTGGTGA
GGTGTCTCTACCCGATTCTGGCCCCGTCAGGTGAGTGAAGTCTGTAAGCTGGGG
TTGAGCAGGTGACGGGCTACCCACACAGCAGCAGAGGCTGTGGGGCCCCCTGAGAGGC
GCTCCAGGTACCTCTCTAGGGGGCTGAGCCCGGGGTGACCCGGGACCTCGCTGCC
CAAAGCCGGCGCCCTCTCCCGCCGCCCCGACAGGGCCAGAGAAGCAGGTGTGGGGCG
CACAAACCAAGTCAAGTTCAGATCTCTGTGGGGCGCGTTGAAACTCGAAGCCCCAG
GCTGGGAGGTCTAGACACCCCTGCCAGACCCAGCAGCTGGGCTGAGCTACAGCTGCCT
GGGGGCCAGGGTGCACCTGCCCTGTGGTGGGGTCAAGGGCAGGGAACCTCCGGGA
AGGTCCCCAGGGTCAAGGTGGGGCTAAGCTCCGGTGACCTCTGGGAAGTCTGGGGCTG
GGTTTGTCTCCAGAGGAGAGAGGGCCACTAGCCTCAGAGGGGCTGTGGC/ACGGTGGGA
GGCCCCAGGTGACUCCAGAGCGTGCAAGCAAGCCCCCTTACTGCAAGC
GCAAGGGCAGAGGTGGGGTGGGAGCTCGACCCCGAGCCAGGTACACAGGGGGAAG
GGCGAGGGATCCGGCAGGGGCCACACCCGCCACCCAGGCAGCCACAAAGCTTTGGGC
CCGGAGCCCCAGATGGGCCAGCCAGCTCTGGGAACAGTCTTCCAGAAATCCCCAGCT
CTGGGTACCAACAGGGCTGCCCGGCCCGAGAGCCCTCGGCGGGAGACCTTCCCCAGG
GGGATCTCTAAETGGCAAGGCTTGTGGGAGGGGCTGGTGAGAGGCCACTCTGGCGGGA
AGACCCCGAGCCACCTGGAGCCCTAGCCACTGCCTGCTCGGCTCCCTAGGGATCCAGG
GCCATCAGAGAAGCTCCAGCGACACTGTTTATTTCAAATGACACTTTTAAAGAAAACA
GCCTCACCAATGCTTGGCCCTGAGTCTGGAATGTGCAGACAGACAGTGGCCCTCCCG
AGAGCTTGCACGGCCCTCCGGGTGGGGAGGAGCAGGGGGCACCCCTGGGACCGGGCGG
AGGCTGTGAGGGCACGAACTGTCTCTGGGCCCTGTCTCAATTCGGGTGCCAGTGG
CCCCAACTTCCAGCAGACCCAGCAGGGCCCCAGCTTGTCTTGGCTGGCCGCTGGTCTCT
GTCACCCAGGCTTGAGTTCTGGAAGATTCTGCTCTGCTCCCGTGTGCATACCACT
CCCCGGGCGAGCCCTGCACTTCTGTTCTGCTGGGCTCCCTGCCTGCATCCGTGAGGCT
GCAGCCCGCTGATCTTCCAGGTCTCTCCGAGCCCCCGCTCCAGGAAGCCCTCCAGG
AGAGCTCAGGAGGGTGGCTCCCTCGCGCAGCTGTGAGACCCCTGGGGCCACCCCGCG
GCTGTAGGGTCCAGGTTCCTCCACAGCCCTCGGGCAGAGGCTGGGGCGCTGGGTCCCTC
GGAGACAACCTGGCTCCGAGGCTTGGCTAGACGGGTTCGGGAGCCCGTCCCCAGCGG

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FIGURE 8, CONT'D.

CACCCACTGAGTTTGAACACTTGGCGCCACCCCCACACCCAGGCGGTGGCCAGGAGGC
CTCCTGGGCAGCAGACAGTCCGTGAGGTGGCCCTGGGGTGGCTCCTGACCTGGGCGCTGG
CCCAGCCCTGGGCACAGCTTCCAGATCTTGCTGCGGCTCCTCCAGGCTGCTCGGCC
CCTCCCGCCTGGGGTGGCCAGCTTTCTGGAGGATSCCCACCCCTTGCCCATGGTCAGG
GAGGGGCTGAGAAACCCACCTCCTGCTCTGCCCAGGCTATGCCAGGGGAACAGGTTT
CCTCCCGCAGGAGGGGACCGAGTCCCTGACAGCCCACTGAGAGGGGAGGAGGTGCTGG
CTCTGCCCCCAGCCCCACCAACCCGCTGGCTTCTGTTCGAGCCCCACAAGCACTAAA
GGCCCCAGGTCTGGAACATCAAGACCCGGGAAGTCCATTGTATGAATTGAGTGTAAA
TGAGCCTGAGGCTGTGGCTTGCTTTCCCACAATTACCGCTGCCCGGAAGGGCTCCGG
TACCGACACAGCCCCAGGGGCCCTTGCCCATGTGGGAGCCAGGCTGGCTGAAGAAG
CCCCATAAGGTGGACCCCACTTTGAGCCCCACGAGAGTGGGCCAAGGACCAGGTGAGG
GCTGCCAGGCTCTGGGCTCCTCTGCTGCCAGGTGGGCTCCTCGGGGCCAGGCTGG
CTTGAGGACCTTCCACGCTGAGTTCCCGAGCCTCGTATGAGCGTAGTGAGCGGAGCC
ATGCCAGCACTCAGGGGCTGAGGGACAGAGCGGGAACCTCCAGCCCCGGTCTCGGC
CCCTAGGATCTCTTAGGTGGGGAGGCCAAGGGAGCAGAGGGGTGAACGAGCTGTGTG
GGGCCCCAGGCTGCCAGCAGACCCCTCCTGCTCCAATCCTCGGCCGAGTGGGCGCCAG
ATGCCGGGGCAGTGCCATTTCCAGGCCACCGGAGGCTCCACAGGGAGTGAGGACAG
AGCTGGGAGGGAGGGCGGGGGGGTGGGGAGGACAGAGCGGAGGCGGAGGCGGCTGAG
GAGGCCCGAGGGGGCTGGAGTCAATGACCCAGGGATTATCGTGTGGGTCTTTGCAAA
GTTGGCTGAGCAACGCGGAGCCAGGGTTCAGGGAGAGGGGACTGGCGGGGCCCGCGG
CCCCCTTTCCCTTTCTGGAAGAGCTGTTCAGAGTCAAAATCCAGCTCATGATCCG
CCCCCTTTGGGACTGATGTTTCAGAGGCCAGTGGTCCAGCACTCTGTCCACCGCCCG
CCACGCTCCCGGGGCCGCCAACCTGTGGGCTGCGAGGTGCGGGCACCTCTCCCTTCG
AAGCAAGCCCTGCCCTGCGTGGGACGCTGATTTCTGCTTCTCTGGGCTGCACTTTG
ACTGGGGTGGGGGGTGG

Contig 4 (1592 bp)

AGCCCTCAGCCCTCCGAGCAGCTGCTGGCTCAGCGGCTCGCCCCGATGTGCGGC
CTTCCATAATCAATCATGGAGGGCCGGGCGGGGGGGGGCGGGCCGACCTGTGAGCCAGC
TCCAAGGCGAGGACAGCTCTGTTCGGAGGGTTCCAGGGGTCAGCCCCACAGACAG
CGGCTCGGCCCTTCCCGAGGGGACCCCCACGGAGGGCCAGACCGGAGGCACTC
GGGGCCAGAGGCCAGGCAAGAGTGAAGGACGCGCGGTGGGAGCGGCGGTGAGCGGG
TCCAGGCTTCACTTCCCAAGGAGCCCATGCCCCGAGCCCGCACTGAGCCCTGTGACGCC
TGTGGTGCCCGCAGGCGCCGACCCCGCCCCCACCAGCTGGGGTGAAGGAGGGAG
GGGGTGGCTGACCGATGGTAACAGCTGCTCCCCCACTCGCGGGCTGGACAGGGCTC
GCTTCTCTGCGCGAGCCCGGGCTGCCCCATCCGTACGGCCACCCAGGACTGTGCGT
CCAGCCTCCCTCCCTCAATCCCCCGCATTTTCGAATCTCGGGCACTGCTGCTTC
CTCTCAAATTCTGGCCCCCTCGCCCCATCCCCGCAATGGGAAAGGCCCGGATGCCA
GGACACTTCTCTCTCGGCCGGCGGGGGAGGAGCAGCTGGCTGGGCGCCGCACTGT
GAGGTGGGGGCTGCCAGGAGAGGGCCAGATTAGGGGGCTCATGGGAAGCTGGGA
GGGAACGCTACCCAGAGCCCTCCTCCCGAGCCTGTGCTGCTCCTCTCCGCAATTTCTG
GCCTCTGAGTGTCTCTGGAGGGAAGGACCACTGTGCTCTCGCGGCTCTGGCTCTGCC
AGGAATGTCATCTGTCCGGGCGGGTTACCTGGCTCAGAGCGTGGTACCAGCTCATCC
AGCCCTGAGGCTGTCTCTCGGGAACAGTGGATGGGCCAGGCGCCCCCGTCAACCCCGCA
GCTGGGCTCCACAGACGGGCCCGGATGGCCACGGAGGTGGGGGCGGCCCCAGGGCGAG
GCTCCCTCTGGAAGGGCTAGAGTGTGGGCTGCGCGGAGAGGGAGGCGGACGCCAGGC
CAGGTGACGCCCGGGCAGGTGCTGGTGGGGCTGTGACCCACGTGTGAGCTCAAGGGT
CCAGGAGCCCCAGGACAGAGCTCAGGACAGACCTCAGAGCCACAGCAGGAAGCCTG
GTGGCAGTAGCTGGCGGGCGGTGGGCTGCTCGGCCCTGCAGACAGAGGACAGGCGAGG
TCCCTGCTGATGACAGGGCTTTCTGTCCCTGGGGGCGGAGGGGGCCGACCATGG
ACCCCGGCCCTCCTCTCGCAGGATCCAGGCGAGCTGCTCAGGCACTCAAGGTG
CACAATGGTCTCCATCGTCCAGATTGACAGGCGAGCACTCTCCACTGGACGGCGGCC
GGGGTGGCTGCACCGCGCTCAGGGCTCAGGGCGGGCGGCCAGCCNCCGAGGCC
TTGACCTGTCTTATACATCTCAACCTG

Contig 5 (831 bp)

TGAGATGTGTATAAGACACAGGCCTTGACCTGGGCTGGCTCAGCTGCGCGCCCTCCTC
CTTGACGCTCCGCTCGACCCCATCATCAGCCATTTCTACCCCTTCTGTAATAAAAA
ACCCGAGGCGCGGTGGCCCCCTGTCCGCTGGGGTGAAGTGGGCTGCTGCTGGTGGCTC
CCACCTGGGGCGGGGCCCTGAAACACACACCCGGGATGGCTTGGCCGGGGCCCTGGT
GGAGGGCGGGGGGCTCGCTGCTCTGTGCTGAAATTTCCGTCCACATGCCCCGAC
TCTCTCCCGGCCACCTGACGGCCGGCGGTGCCCCGCCACTTTCCGAAGGACGG

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FIGURE 8, CONTD.

ACTCAGCATTTCCAGGGCACCTGCTGATGGTGCCAGACCCCGGGGCTTCCCGCCGG
GGCGGGCCCCACGTGCCCCCTCCAGTGGCCACAGCGGGCTGGGCCAAGGCTGGGAGTTC
TGCACGGGCTGGGGGAGGAAGCGGGGAGAGGGGACAGTCTCTGGCGGGACGAGGG
TGGGGGACGAGGTGGGGAGTTCACACAGCCGGGGACGGGGACGCCGCTTGGCTGCCCT
GGGTCTCAGCCGGGGACACTGCCACAGGAGAGAGACGGCAGACAGTACAGCCACCCG
TTTTATATCTCTCAGCGGCTGTGCTTTATTGGGGTAAATATGCAGGACATAGAAACT
CTGCCACTGGACCCCTTGGCCGGGGACACAGCAGCGGCTTGCATGCTTTCTGGGTGCA
GCGCAGCCAGCACCCCGGCCAGAGACCCCATCTTCCCGATCACCGGAC

Contig 6 (4634 bp)

CTCTGGGCTAGCACCGTGGGGCTTTGCCAGAGTGGAACTGAAGTGGTCCACCCCGGAG
CCCAGAGGGCGGTGAATGGGAGGCAGAGCCATCCTGGGAATGGACCAAGAAAGGGAG
CGGGGGTGGGGGAGGGGCATCAGATCCTGGTCTTCTTGTGCGCTGCGGTCCCTCTGC
CACCCTCCCCAAGCTGATCTGGAGCACACGCGTCGTTAAAGCCGCTATCGAGGCCCCA
CTTCTCACAGACGGAAGGGGCGAGAGTGCCTTCTCACCCTGCTGCGCTGGGAAGGGCC
CTCCCTGCAGCCGAGGAAGCCAGCAGCAGGTGACAGAGCCAGGGGCCAGGGCCCGAGGG
AGGGGCTCGCGGCTCCGAGCCGGGGTCCCTTGGCGTCCCCATCTCTCGTCTCGAGCC
CTCTGGGTGACACAGGAATGTCAAGGGCGGACGCCGGTGGGCGCGGGAGGCGGGTG
GGAGGCGGGCGGGTGGCTCTTACGGGCGGGCTGAGAGATGGGCGCCCTCCGGCC
TGGCGTCTATCTCTCCGCTCTTACCCACTGAGCAAGACACAGAAATGAAGCTCGAA
CGAGCACAGCCAAAGAACGGCCGTTTCTGCTCTTCTTCTTAATCCCTTGGCTTAGGGT
TTCGCGGCTGGACAGCTGCCAAGGGACATGGGCTCCGTCGCGGACATTACAGCA
GTGACCAATCCAGGCCACCCAGGCTGTGCCCTGCGTCTGGGCGCATTTCCAGCCGGCC
AGAGATGGAGCAGCCACTGCGGGTCCCCAGTCTCGGTGAGACAGTCAAGGATGGACCT
GGATGGAGACCGCGCTGCGGCCATGTCCGTGGGTGAAGGAGGCGTGCAGGCGGTGCTGG
GGACATGGTGTCTGCTCCCTCGGCCAAACCATGAAAGCAGCCCTCTCCCAACCCCA
GCACCAACCCGAGACCCCTCGGCCGGAGCCAGCACGGCCACCGTCAGCTCTCGGT
GTCCAGCTTGGGACAGGTGAGTTCAGATGTCCAGGCTGGAGCTGGTCTTGAAGATCC
TAGGGGTCCAGCCACGACAGGAGGGCCAGGTGAGAGCCCCCTGTGGTCTTAAGGATGCA
ACCAGGGGCGCGGGGTGCTTCCCTAGAGGGGTAACTCGGCCCTTGGGACCACTC
ACCCAGGAGGTCCAGAGCCAGCTCGGAGGGCCACAGGTGCCAGAGTCCCACTTGG
GGAAAGCTGCCCTCTGCCAGCCCCGAGCCGGGCCCTTGGCGCCCGCTCCAGCCCGG
ACCCCGGGAGATATTACCCCTTGCCTCGTGAATCAGGAGGCCCGAGCCCATGTTT
CAGTCTTTCTTCCATCCAGCCCCCAGGAGAGAGGTGCTGAATGGGTCCCTTGG
AGGCTCCTGAGCCCCAGAACAGTGCCTCTGAGCAGACGGGCACTCTCAGACCACTCAG
GCTGGACAAGTCACTCTGCTTGCCTGCTGATGGGCGCTTGGGAGAGCAGACATGCTG
AGGAAAGGCCCTGTGCCCTTCACTTAATCCCAAGCCCAAGTCCCACTGGGTGGC
AGCTTCAACCTAAGCAATAATCTGCTCTTAAACAAACGCGCGGAATCCCACTGC
CCTTCCCCCGCCCGCCCCC
ACCCCTGGCTTGACCTCCAAAGCACTTGGAGGGGCTTTCTCAGACACCTCCAACCC
CGACCCCATGAAGAAGGGGTGATGSGGTGTACCCTCAACAGCAAGAGAAGCAAGCCCA
GAGAGGAGTTGGCTGGACAGCAGGGGTAGGCCCTTTGCCCGAGGGCAGGGCTGGTG
CCACCTGGGTGACGGCGGAGGCCCTGGAAAGCACCGGAAATGAGCACACCTGGGTCTCT
AGAAGGTCTCTCAGACCTCTGGGGCTGAGTCATTTCAACACTCCTGGGCGGGCAGGG
CTTCTTCTTGGCCCGAGGGACAAGTCCCTTCTGTCGGGGGGTACGGCCCTGGACCC
CTGTCCCCCGCACCCCACTCTCGCTGCTGAGGGCCGCGGCCAGCTCTGGACACAGATC
CCTCAGAGCCCTTCTCTCTCTCTGCTCTCTCTTCCCAAGATGCCCGGCCCTCAGG
TGGGGCAGCCAGGCGGCAAGATGTGTTCCAGGCTCTCGGCCCAACCCACACCCCTGC
TCTGCCCTGACAGCTCCAGAGCAGGCACTGCTGCTGCTCTGCTCTCTCTCTCTCA
TGGCAGAAACGGTGCCTCCCTAGCTTCCCCAGAGAAGGGAGATCTGCTCTCCCGGAG
GACCTGCTCTGCTGTCTCTCGCCCGGCTTACGGGCTCTCCCCAAGGGTGGCCGCG
AGGAGGGCTCTGCTCCGGCCAGGGGGCTCCATCTCCCGAGCCGACAGGCTCCGCT
TGGTGGTCCGACCTTCTCCCAAGGCCCGCCCATCTCTCTGCGCTCCCCAAACCTG
CCTCTTCCCCAGCGCTTGTCCACGGAAGACCTCCACCGTGCCATTACAGCTC
TGGCCCCACCTCCAGCCACCCCTTCTCCCTCTCTGGAAGCTCCCACTTCTCTC
CGCTCTCCACGGCAGCAGAGGTGACAGCTCAGGGTCTTGGGGCTGGAGATGGCC
TGCCCGGGGTCTGCTGACCGCTCTACGGAAGCTGTGCCGGGGGTGGGGGTGCTCT
TGCCGAAACGGTGGAGGACGAGCCATCCAGGGCAGCCGGAACCTGCTCTTGGTCT
GAGACGGAGAGGCTGGGTGACAGTGGCTGAGGGGCTGCACACAGCTTGGCTTGGGTCC
CTAGGTGACACACTGCTGAACATCTATTGCTGCTCCCTTCCAGGGTACCTTGGG
TCCCGTGTGGCTCAGGGCACAGGGGGCCCAACAGGCTTACAGAACCCCACTGGG
ACTGACCCAGGGCCACAGAACTGGGGGGCTGGGGTCCAGAAACACCCCAACA

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FIGURE 8, CONTD.

CAGGCCAAGGTGGCCAAAGCCCTTACTCGAGCGGGGCTGCCCGTCCCAAGAGACTCTSGCC
AGTCGTCCGGATCCAGCTTCCCGGGCCGGGCGCCCGCTGGGCTCCAGGCGGTCTGGG
GGGCCCTCCCGGGGGTTCGCCCTCCGCTCTCAGCAGCAGGAAGAGGAGCGCGGCCAGC
GGATGGGGAGAAGAGGGCGCCCTGGCCATCTTGCTCCCGCTGGGACTTGAGGAGGGTCTC
GGGCCGGGCAAGCGGGACCGGGAGCCACAGAGACCCTGGAGGAGGCAGCATGGCGGGGAG
GTGACCGGGGAAGAGGGCCGTGTCCAGGCTCAGAGCCCGGCTGGCCGCGCCGGCCCTCG
GGAGGCGTGGCGCTGACCGCTGGCGGGAGGTTTGCTGCGTGTGGGGTTTGCAGAAAGT
GCTGAGCTGCTGAGCCACAGGCCAGGCTCAGAGGGACAGGAAGGAGGTTGCTGCCAG
CCTCGGGCACTGCTGACCCATCTCCCGTTTCCAGGGCACCAGAGCCACCTAATCTGCCGG
CTCTGTGCCCAAGGACAGGCTTGCTGATCTCTCAAGGCCGGGCGCTCCGCTTCCCTGG
GAGAGGGTTAAACATCCAGCCCCAGCCAGCATCTCGGCGAGGTTCTGGCTCCCCCGCT
CGTGCTCTCTGAGACCCCTGCTCGGCACACCTTCCCTTGAGAGGAGGAGGAGGAGAA
AGCGGATGGAACCACTGACCTTCAGCCCTGAGGGCACCTTCCACGTGCCCGCGCCG
CCCCGCTCCTCCGCCCCAGTTCTCAGGCCCCAGTCTGTATGAGGAGGGCGACCTC
CGGGCTCCTGGCTCCCGGCGGCTCCGGAAGACAGGGCCGCTCGGCTCGGGCTGCAGGGA
GGGGCCCCAGACGCAGGAGAGCAGCCCGGAGGCAACCCCGGGGTCTTCCAGAAGGAGG
CCTGGCAGGGGAGGGGGGTGCCACCACTGCTGTCTCTCTGTCACAGTGGAGGGTGT
GGGTGGGCAGTGCCCGGGTGGGAAGTGCAAGAACCCCTGGACCTGGGGCTGGGCGCC
ACGGGGAGCGGGTCTGTGAGGACCTGGGGGAGGAGGCGAAGGGCTGGCCAGAGG
CCGGATCACTTCCAGATTGCTGTGGGACCAAGGGCCGACCTCGGGGTGACTTCTTTTG
TGTGCTGGCCACAGGGGGCCCCGGCGAGTTCACACGGAAGGGGCTTCGACCTGGCCT
AACAGCCCACTCCCGAGGAAGATGCAAGGGGAGCCAGACGGAAGGGCCGAAGGGGGCA
TCGGGGACACCGCGGAGGGCCGGGCGAGAGAGGGAGGAGAGGGCAGAGAGGGAGG
CAGAGGCGAGAGAACGGAGGAGAGGGGCCACATGCTTGGAGGGCCAGGAGGAGCGGGA
ACGGCGTCCGGCGTCCAGCGGCAATCAGGCCCTCAGGCGSAGGGTGGTGGACCTGCC
TGGCCTTACGAGCACAGTCAGCAGGCTGTCTTTATACACATCTCAACCATCAT

Contig 7 (482 bp)

AGCAATGGGGCCSTGACCTAAGGAGGAGGCCAGGTGAGTGGGTGACCTCTCGTGGCC
CCGATGTTTGGAAATCCCAAAATCAAAATGACCCATCCGACAAGCTTGATGCTGCAGG
TCGACTTAGAGGATCCCGGGTACCGAGCTCGAATTCGCCCTATAGTGAGTGTATTAC
AATTCAGTGGCGTGTGTTTACACCTGCTGACTGGGAAAACCCCTGGCGTTACCCAACTT
AATCGCCTTGACACATCCCGCTTTCGCGAGCTGGCGTAATACCGAAGAGGCCCGCACC
GATCGCCCTTCCCAACAGTTGCGCAGCCTGAATGGCGAATGGCGCTGATCGGGTATTTT
CTCCTTACCGATCTGTGCGGTATTTACACCGCATATGCTGCACTCTCAGTACAATCTGC
TCTGATGCCGATAGTTAAGCCAGCCCCGACACCCGCCAACCCGCTGACGGCAACCC
TT

FIGURE 9

Human clone af087017.em_hum1: H19 gene + flanking sequences

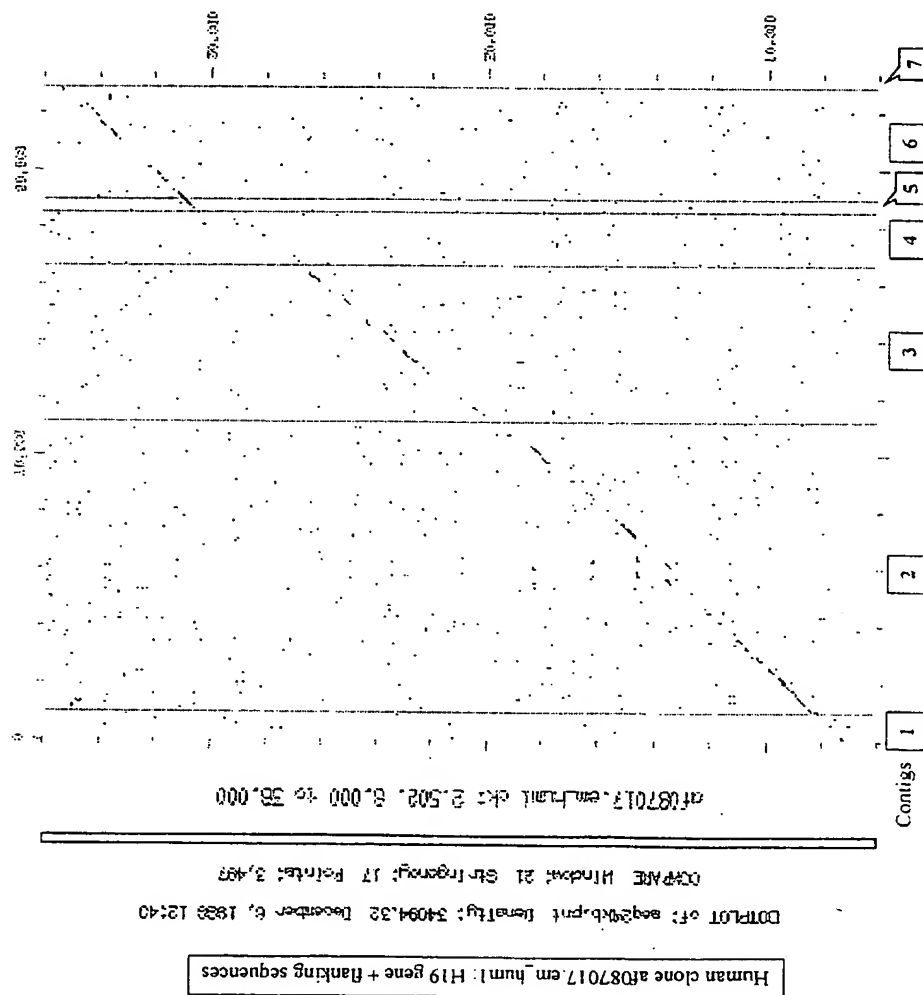


FIGURE 10

IDENTIFIED POLYMORPHISMS:POLYMORPHISMS TYROSINE HYDROXYLASE GENE - CONTIG C3 (figure 6)

1	GGATCCAGCC(A:T)GCAGCC	1081 bp
2	ACAACCCCC(-:C)TCCCACAG	1149 bp
3	TGCGGAGGGG(A:G)GACCTG	1186 bp
4	AGGT(CAAGGCCAGGT:-)CGAGG	1210 bp

POLYMORPHISMS INSULIN-IGF2 - CONTIG C4 (figure 6)

5	CCC(C:A)CCCC(A:C)CGCCGC	438 bp
6	CCC(C:A)CCCC(A:C)CGCCGC	443 bp
7	CGCCGCAGCA(G:A)GCCG	455 bp
8	GCTTATGG(G:A)GCCGGG	503 bp
9	CACGGC(T:C)TC(G:A)GAGCA	525 bp
10	CACGGC(T:C)TC(G:A)GAGCA	528 bp
11	GTCTGC(A:G)GGCAGGTG	571 bp
12	CAAGCCCGG(G:T)CGGTT	636 bp
13	ACCTC(A:G)AGGCCCCCA	710 bp
14	GC(C:T)GGGCCAGCCGC	867 bp
15	ACCAGCTG(C:T)GTTCCC	903 bp
16	GGC(C:G)CTCTGGGCGCC	1148 bp
17	GGGGG(C:T)GTCCCGGA	1305 bp

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FIGURE 10, CONTD.

18	GCGGT (C:T)GGGGGAGTT	1320 bp
19	CGCCC (C:T)GGTCCCGCT	1400 bp
20	TCCC (G:A)TCTGCCGGCC	1519 bp
21	GA (T:A)GCCCCATCCCCC	1547 bp
22	GG (C:T)GGCTGCTGCGGC	1607 bp
23	TGGCTGC (G:A)GTCTGGG	2222 bp

POLYMORPHISMES IN CODING REGION - CONTIG C10 (figure 6)

24	GCGCA (G:T)TGATTGGCA	341 bp
25	CGCCCCCCCC (-:C) (G:C)GG	2247 bp
26	CGCCCCCCCC (-:C) (G:C)GG	2248 bp
27	GCAGCCGGCTC (C:T)TGG	2257 bp
28	GTTGTTG (C:T)TCTGGGA	2413 bp

MICROSATELLITES

29	PIGQTL1: (AT) ¹¹	112 to 133 bp Contig 57
30	PIGQTL2: (GT) ⁸ GCACCGGTGTGCTGTGTAC (GT) ¹⁷	1074 to 1144 bp Contig 95
31	PIGQTL3: (CA) ¹⁹	223 to 260 bp Contig 105

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